

Le Léman: du témoin des plus grands changements climatiques à la bactérie qui résiste aux antibiotiques



Club de plongée de Plan les Ouates, 29 janvier 2018



Tabula Peutingeriana « Romans world »(Talbert, 2010).

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Walter Wildi



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Université de Genève

1. Le Léman c'est



Photo: Anh Dao Le Thi

1. Le Léman c'est

Le plus grand réservoir d'eau douce en Europe Centrale

Sport et détente

Un haut-lieu du tourisme

La nature pure

Le meilleur indicateur environnemental et la mémoire de son histoire

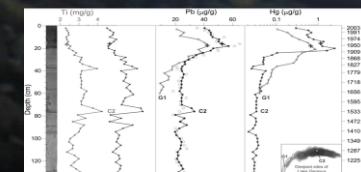


Photo: Anh Dao Le Thi

1. Le Léman c'est

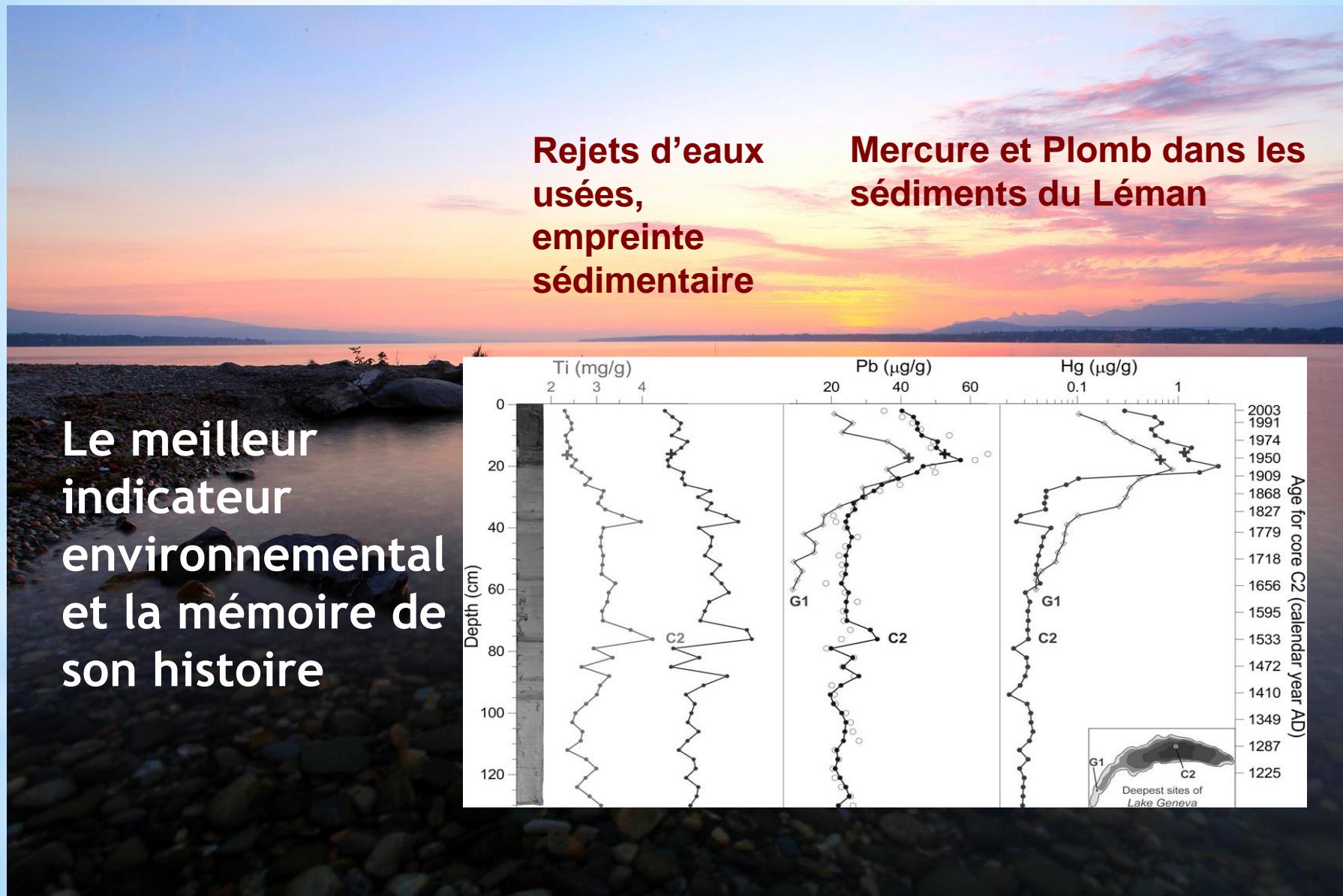
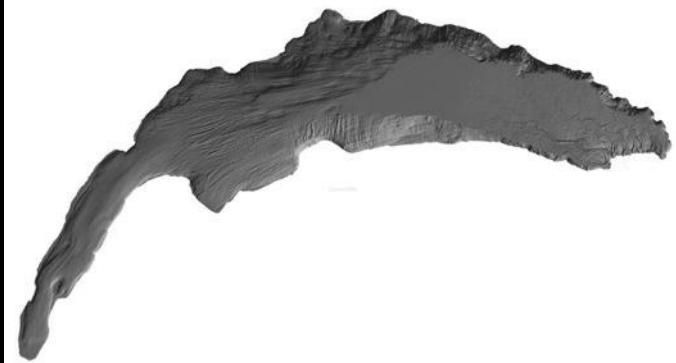


Photo: Anh Dao Le Thi

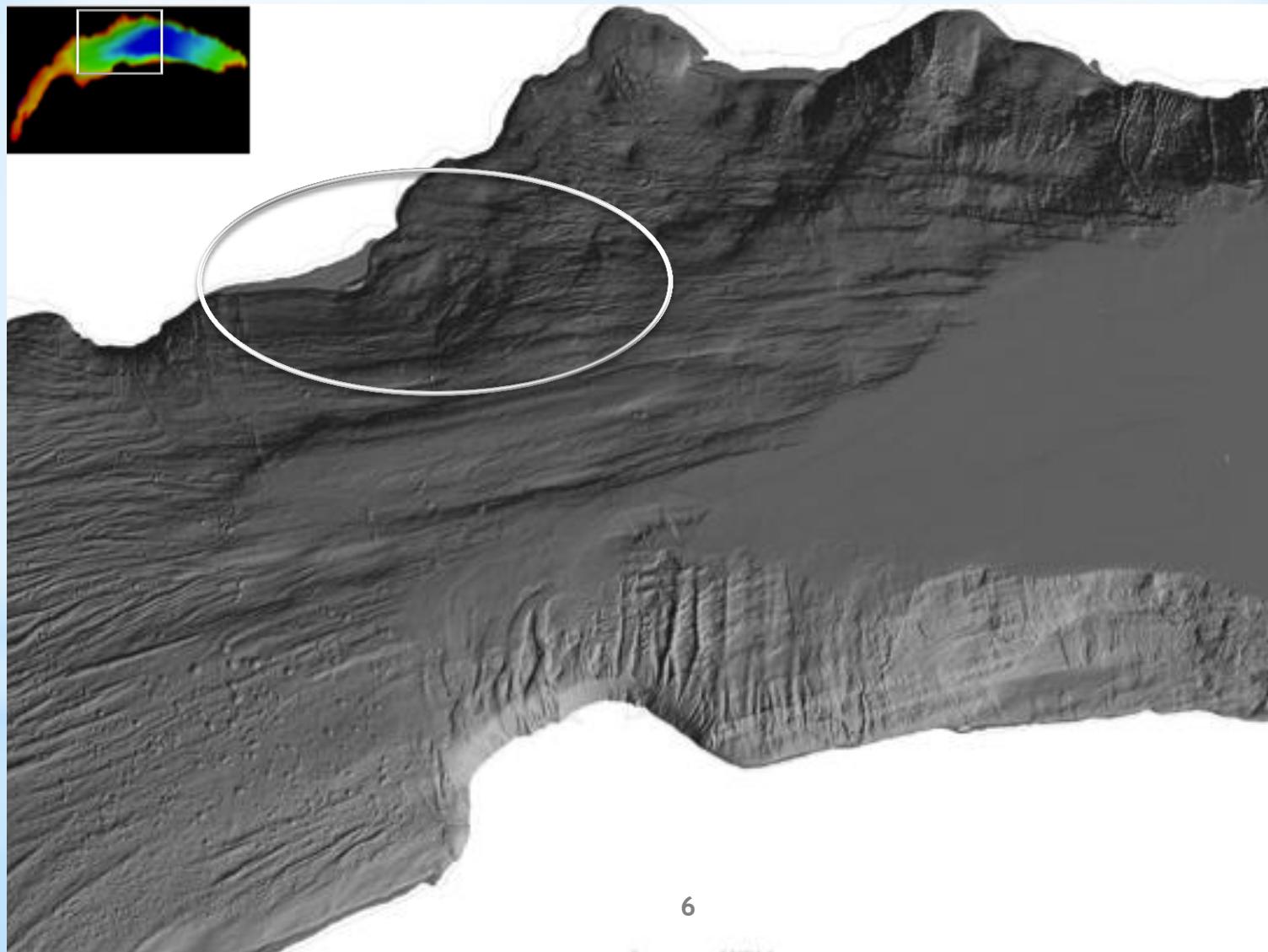
2. Le Léman glaciaire

**La cuvette du
lac: un ancien
bassin glaciaire**

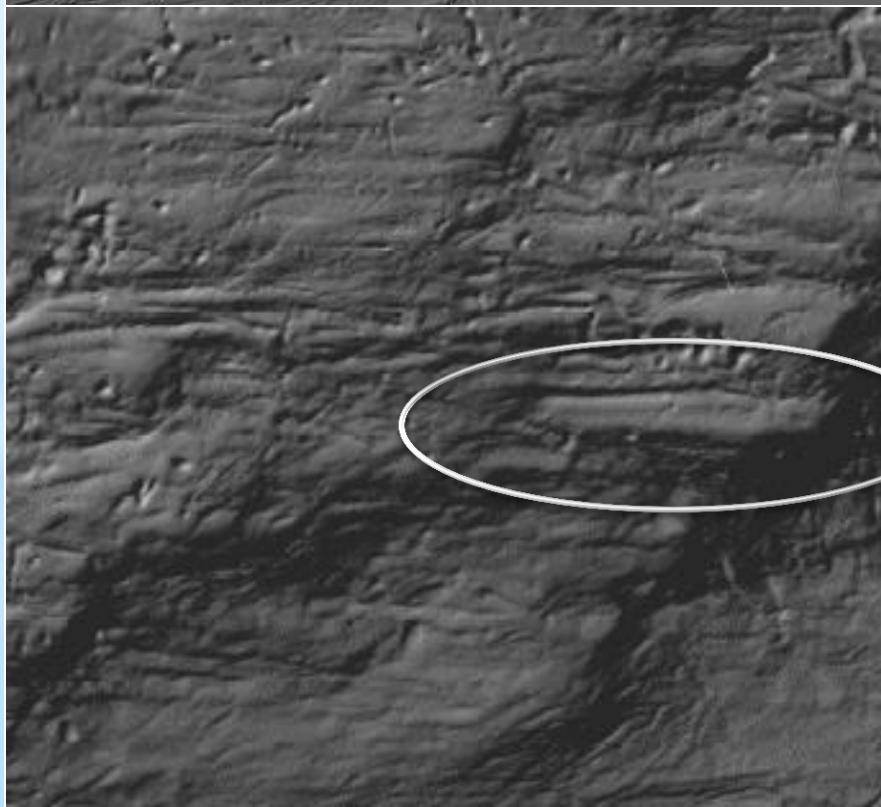
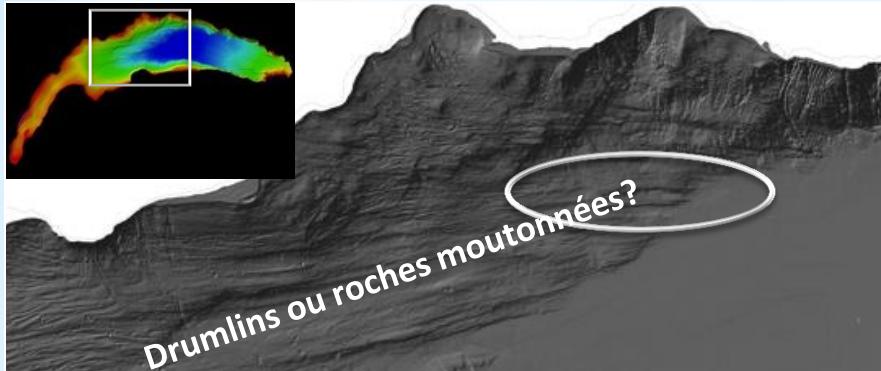


Carte bathymétrique par sonar multifaisceaux (Institut F.A. Forel, UNIGE et Institut de Géologie, UNIBE 2013-2014, Dép. Général de l'environnement, Etat de Vaud).

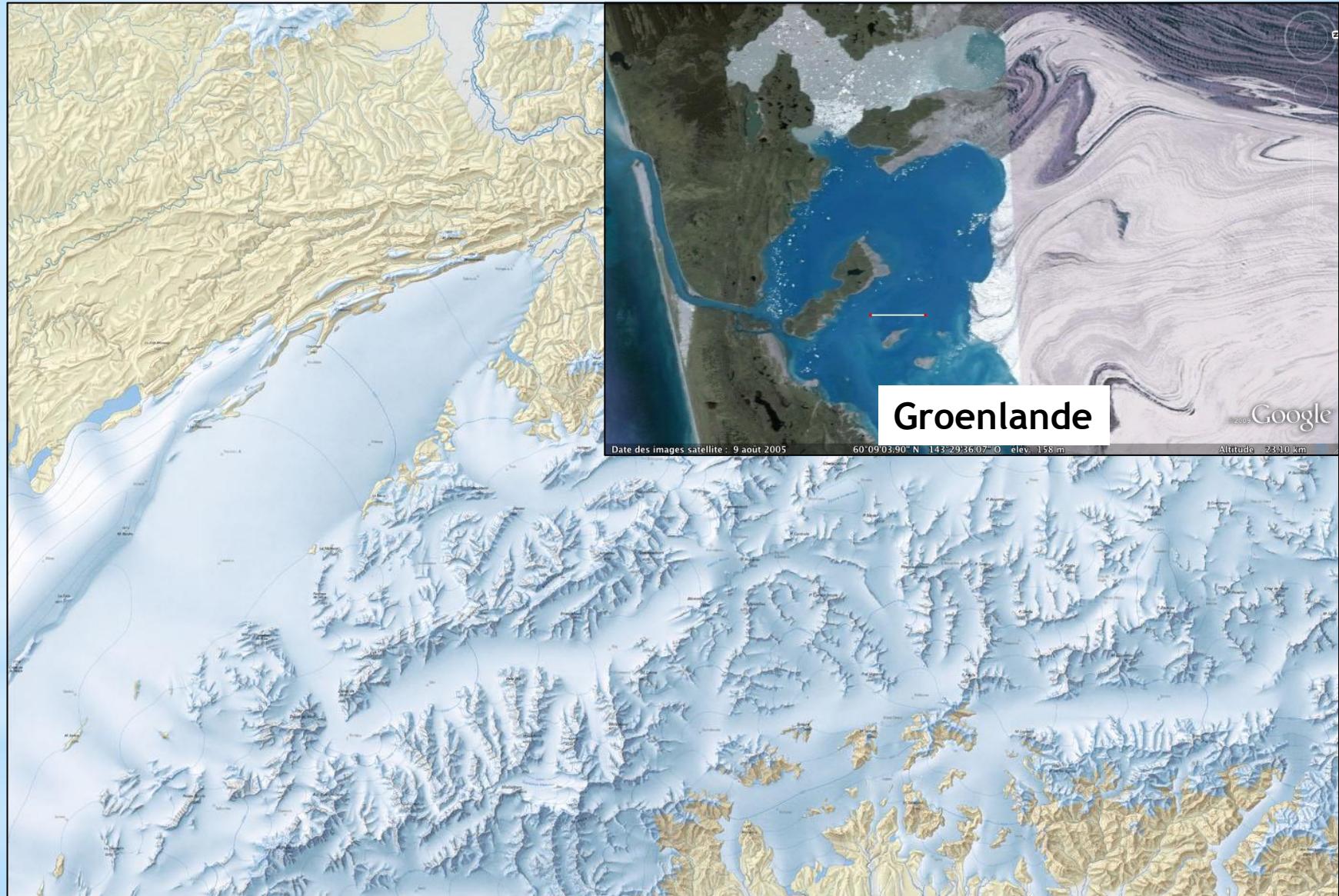
2. Le Léman glaciaire



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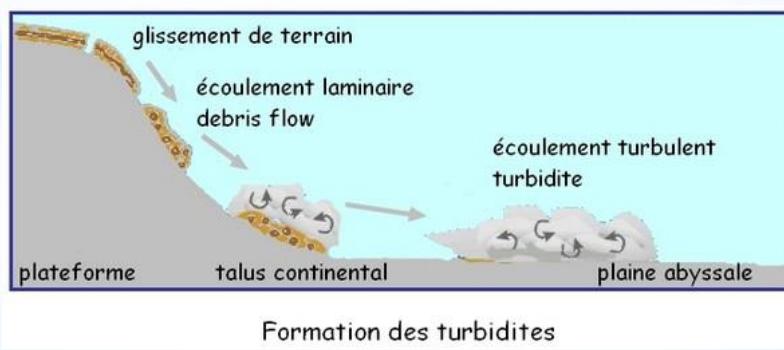
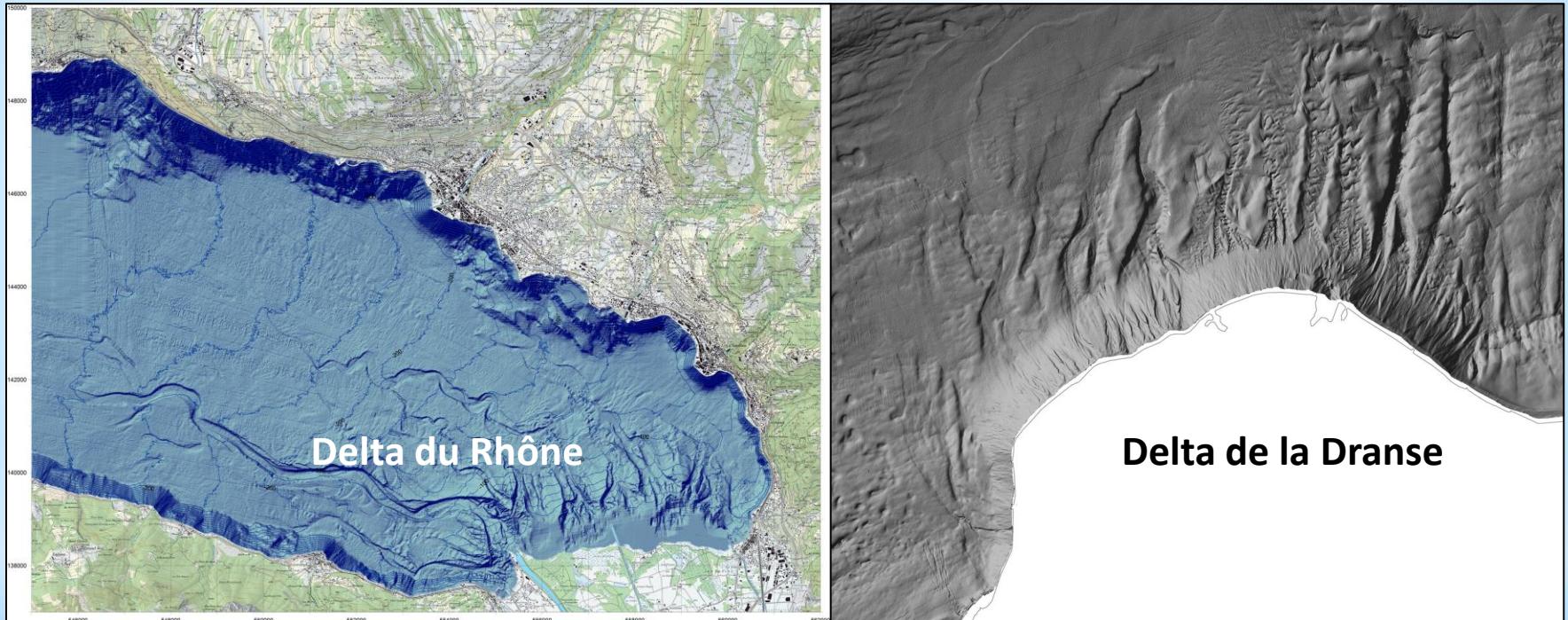
La Suisse au dernier âge glaciaire (swisstopo)

3. Le Léman physique



3. Le Léman physique

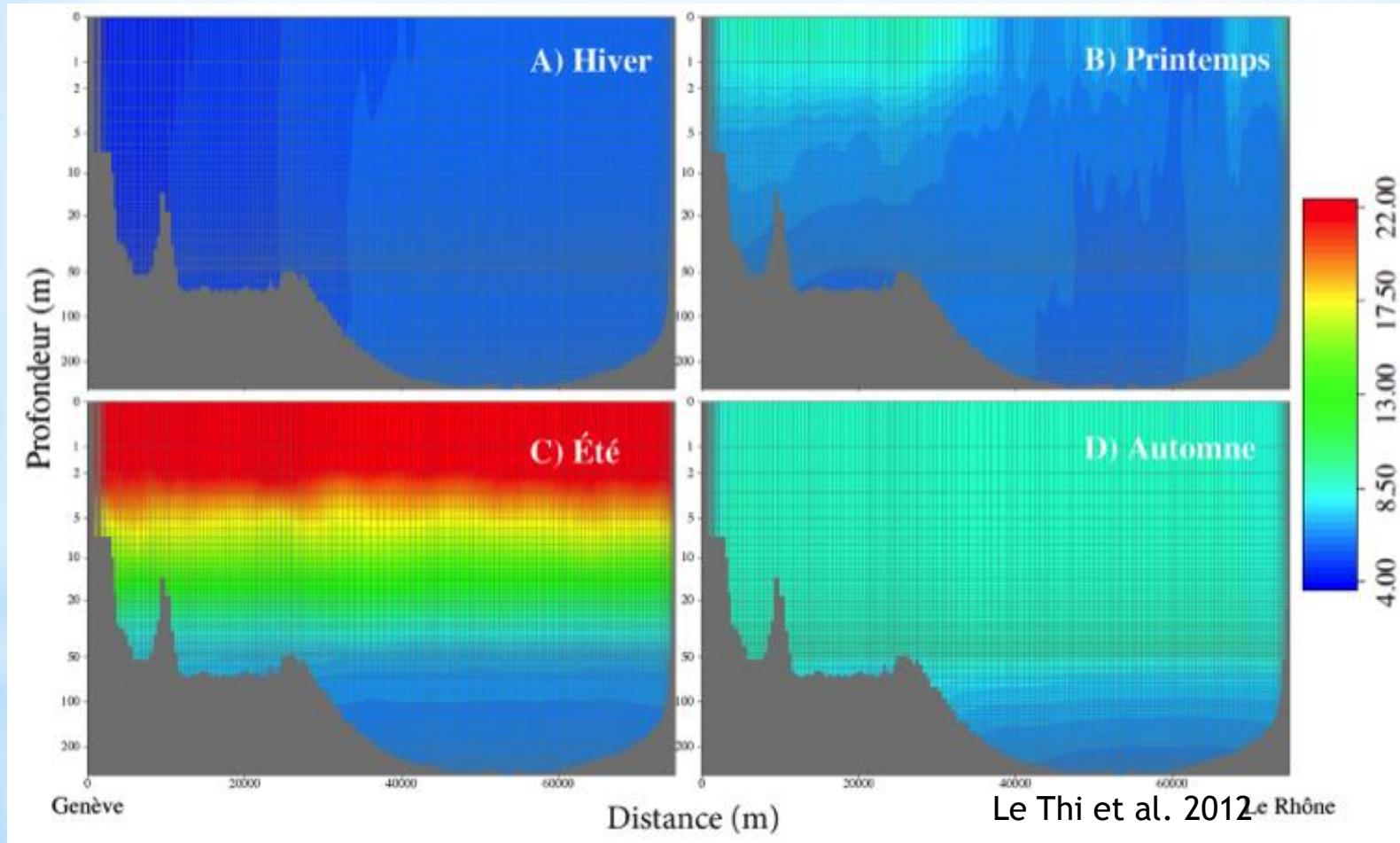
Sédimentation et érosion



3. Le Léman physique

Régime thermique saisonnier (« monomictic »)

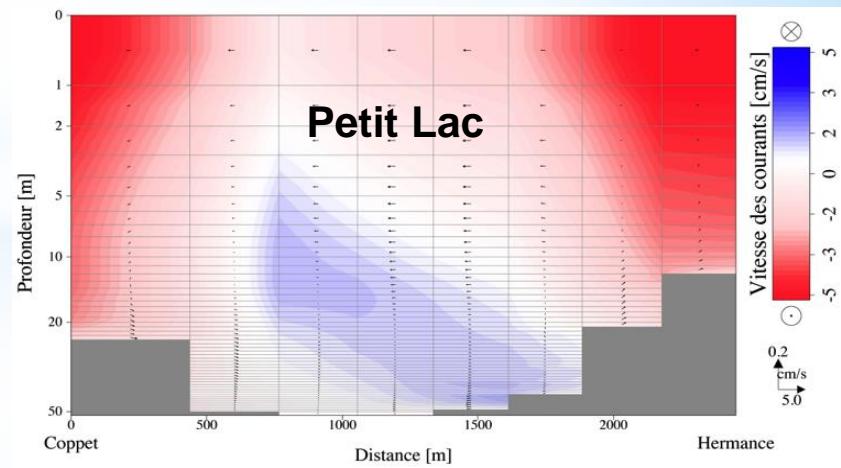
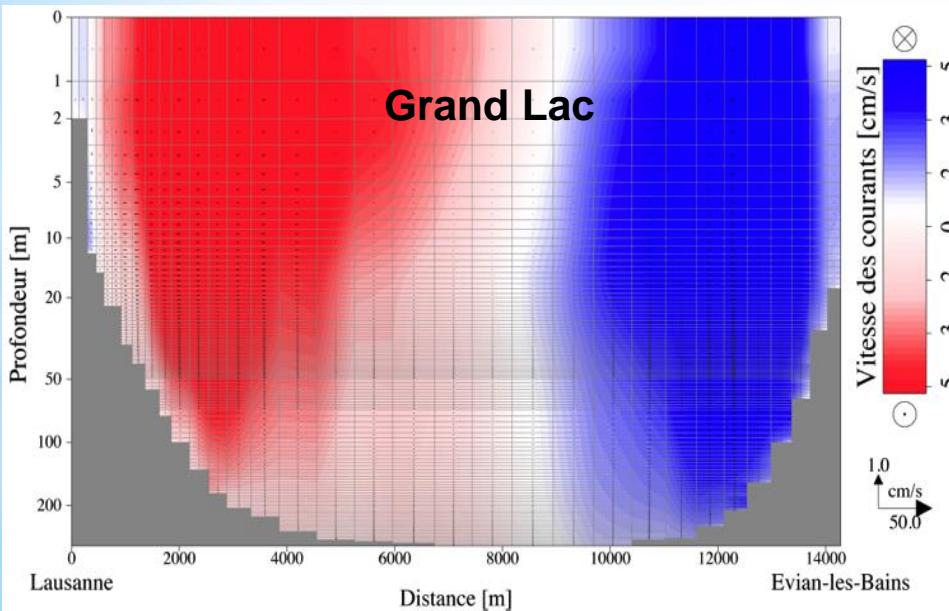
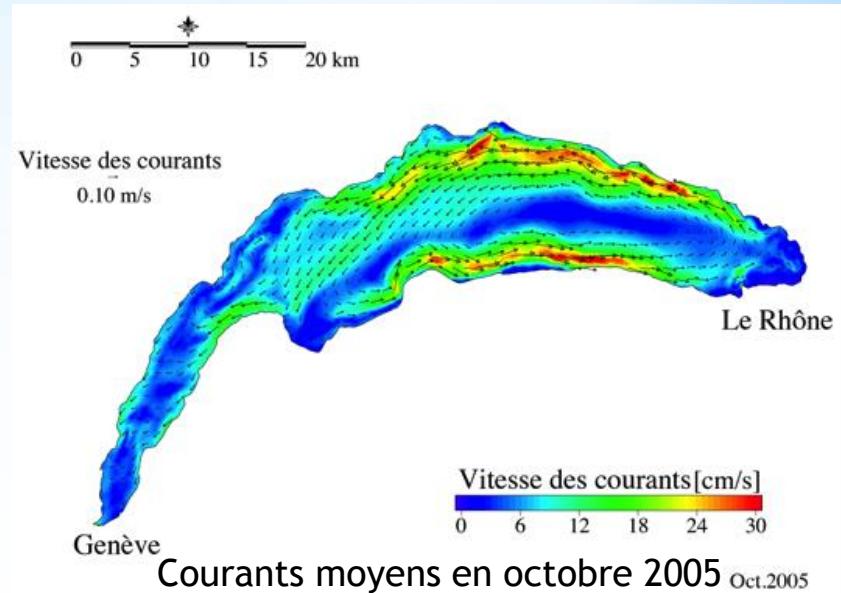
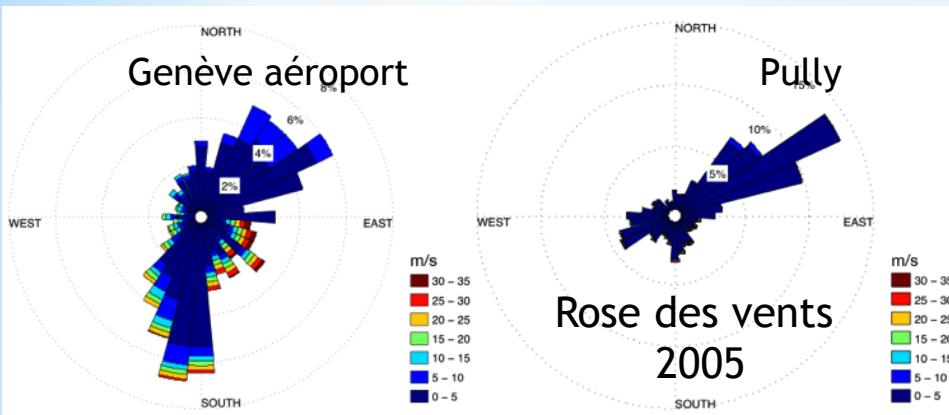
Petit Lac - Grand Lac Lac



Répartition des températures (° C) sur une section longitudinale du Léman
année 2005

3. Le Léman physique

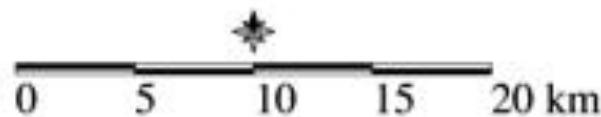
Régime des courants animés par les vents



Courants moyens 2005, transect B, Grand-Lac et Petit-Lac (Le Thi et al. 2012)

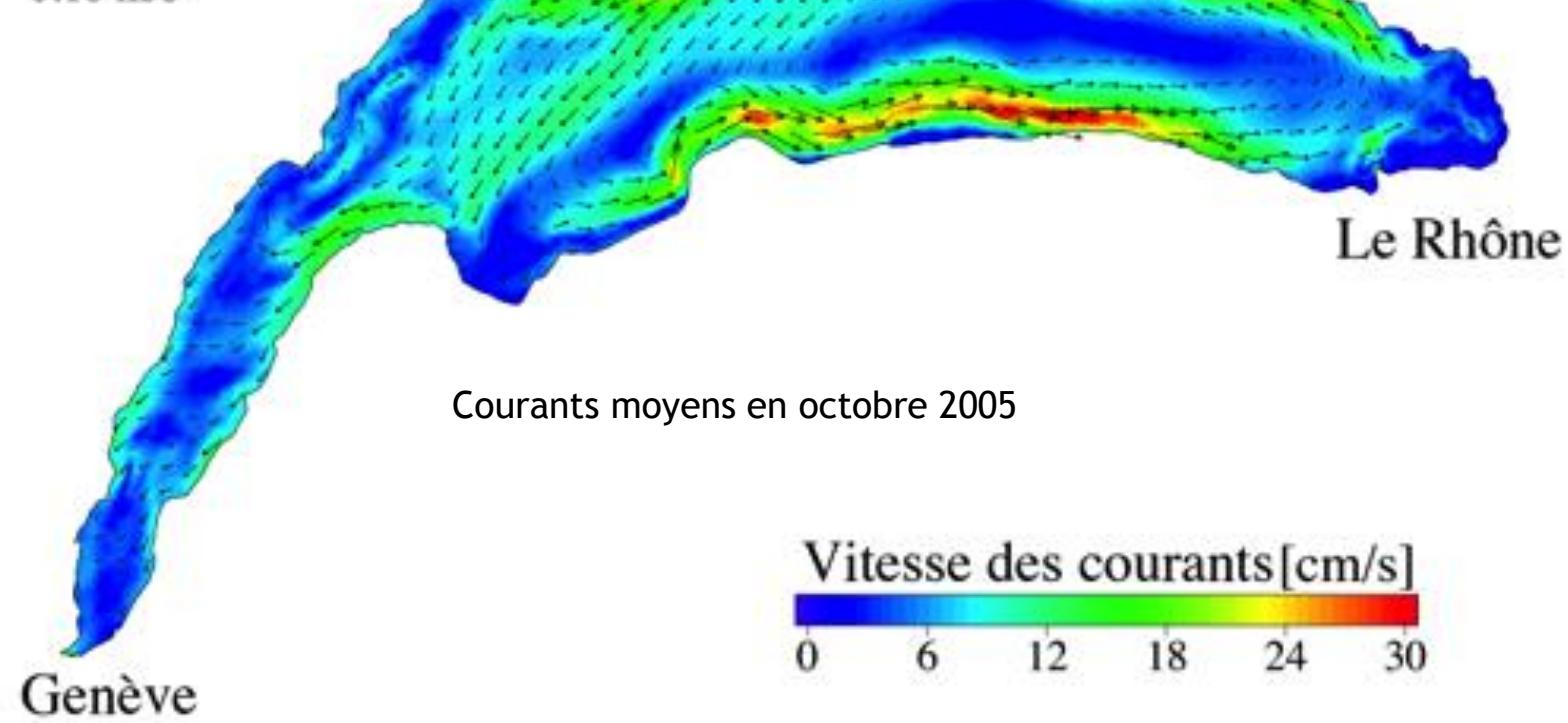
3. Le Léman physique

Régime des courants animés par les vents

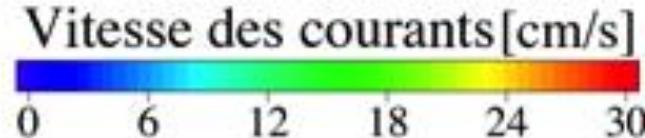


Vitesse des courants

0.10 m/s



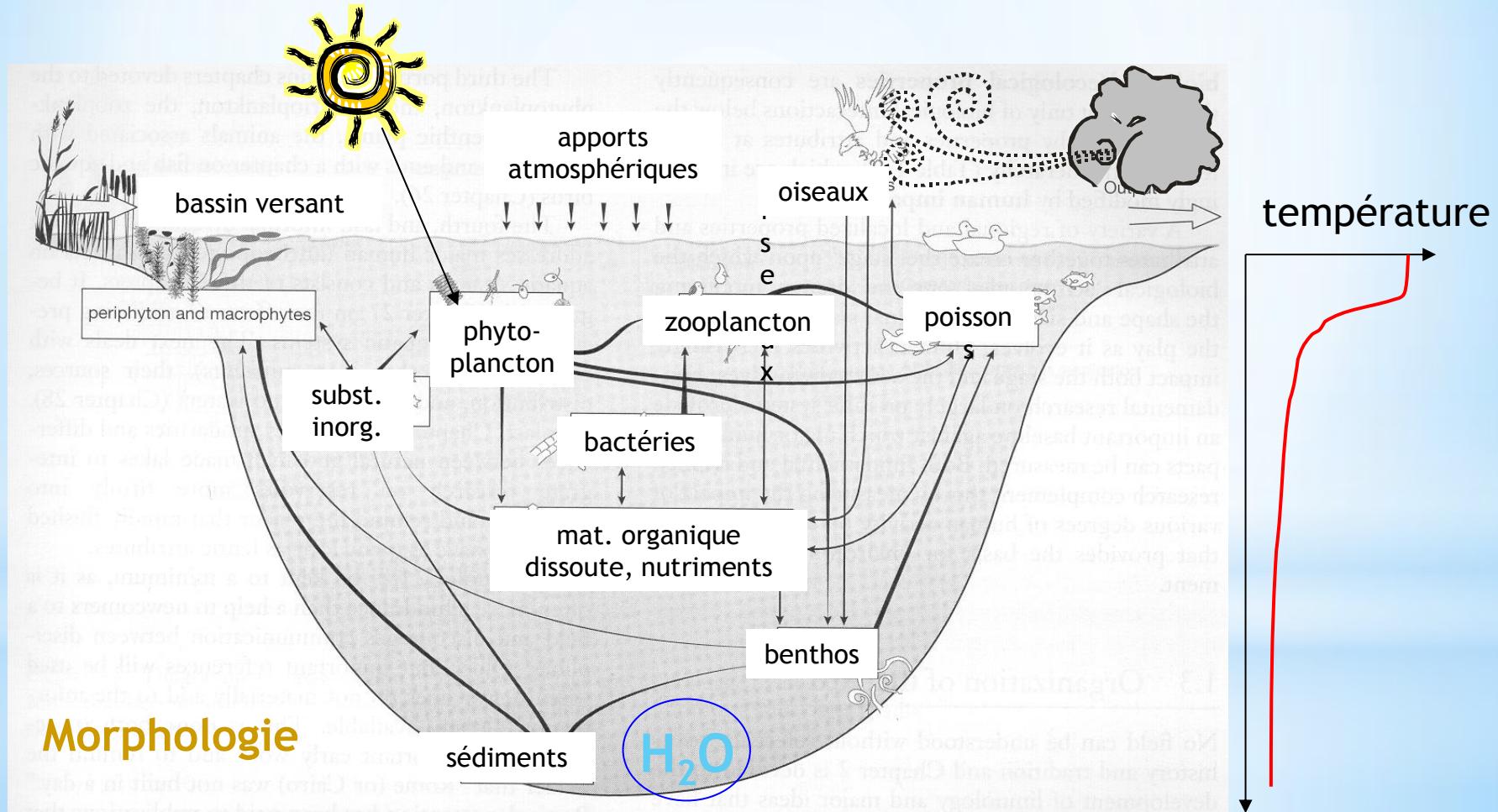
Courants moyens en octobre 2005



Oct.2005

4. Le Léman vivant

Structure et fonctionnement des écosystèmes aquatiques



5. Dissémination des antibiotiques, bactéries et leurs gènes de résistances aux antibiotiques dans le Léman



Objectif de la présentation:

- Les voies de dissémination des antibiotiques en milieux aquatique
- La relation entre antibiotiques et bactéries résistantes
- Comment une bactérie devient-elle résistante et transfert des gènes de résistance entre les bactéries
- L'impact des bactéries résistantes sur la faune et la flore aquatique, comme celle du lac Léman
- Eventuelles solutions pour éviter cette dissémination

Beaucoup de diapos en Anglais

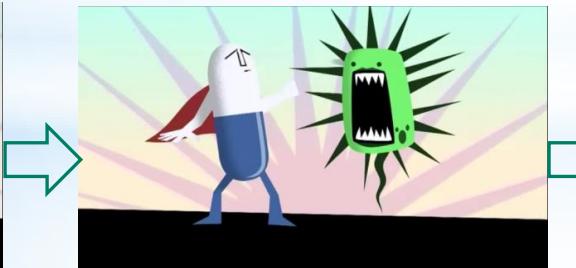
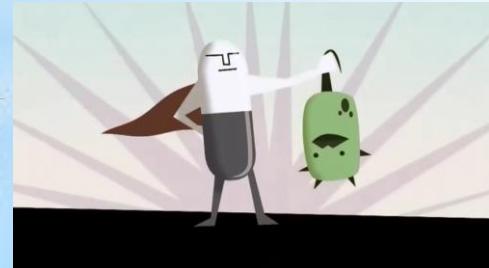
5.1. Antibiotiques et résistances aux antibiotiques



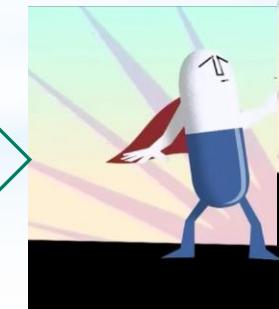
Médicament pour traiter les infections causées par les bactéries: la tuberculoses, les pneumonies, bronchites, otites, méningites, infections urinaires, septicémies, maladies sexuellement transmissibles,.....

What is antibiotic resistance or/and multi resistance?

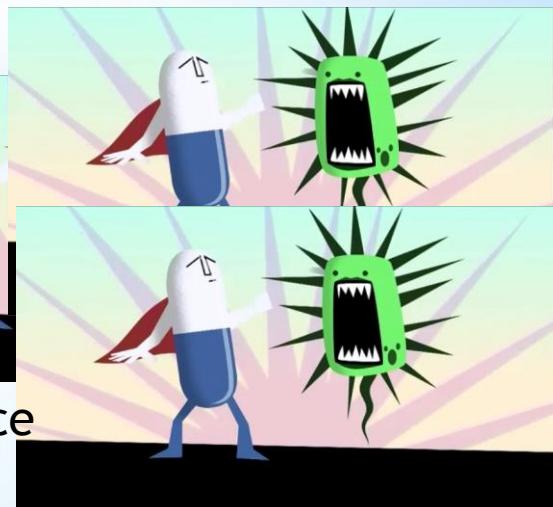
Antibiotic resistance occurs when bacteria develop the ability to survive exposure to antibiotic/s that is/were designed to kill them or stop their growth



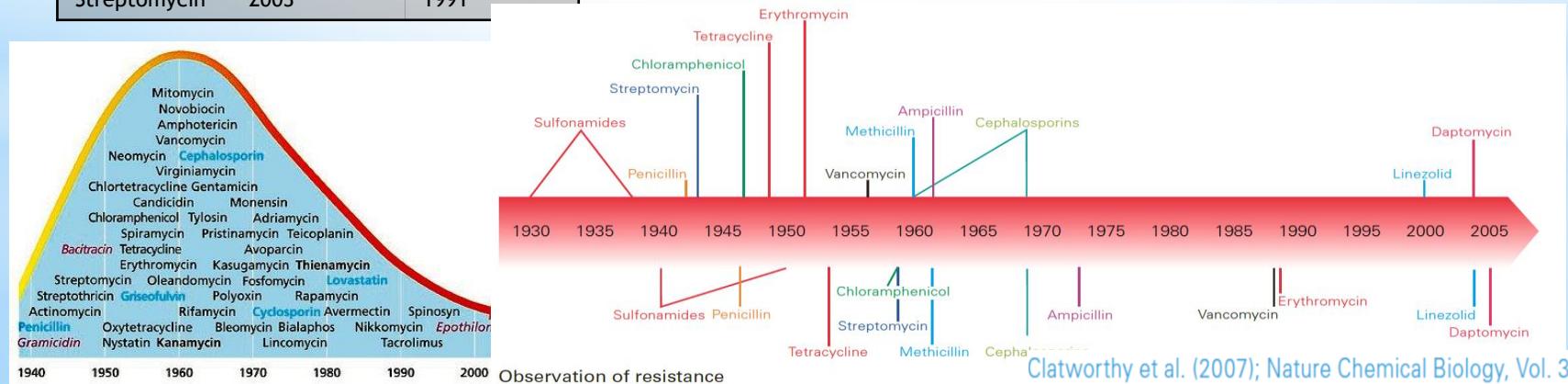
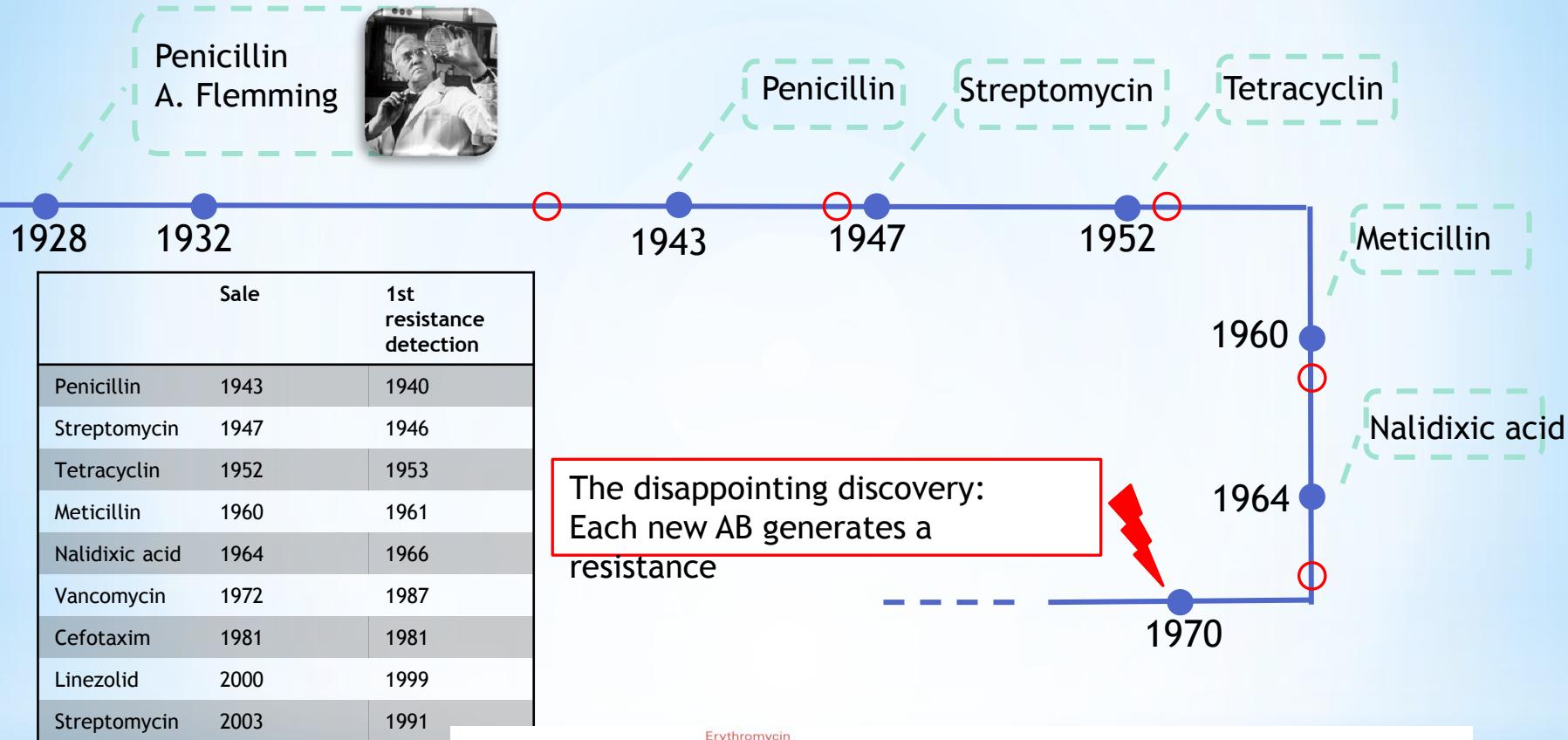
Resistance
ARB & ARGs



multiresistance



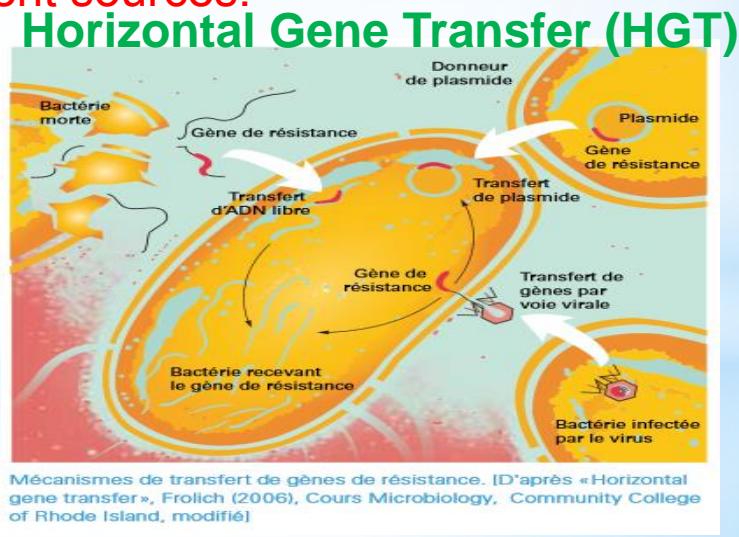
5.2. Quand et comment la bactérie devient-elle résistante



Clatworthy et al. (2007); Nature Chemical Biology, Vol. 3.

Bacteria become resistance by several ways including

- It is important to note that several antibiotics are produced by environmental microorganisms.
- In the recent years the proliferation of multi-resistant pathogens to conventional antibiotics and new antibiotic molecules such the last generation of beta lactamase, carbapenem-resistant and their resistance genes such as (bla_{TEM} , bla_{SHV} , bla_{CTX-M} ; bla_{NDM} , Bla_{VIM} , Bla_{KPC} , Bla_{OXA} and $aadA$). This can be explained by therapeutic application of antibiotic in medicine and veterinary.
- There is the correlation between environmental ATB concentration and bacterial resistance
- Aquatic environment is considered to be of special reservoir of resistance genes since it's recipient of bacteria from different sources.



La prise d'antibiotique/utilisation excessive/abusive/sous-dosage d'antibiotique occasionnant la mutation ou acquisition de gènes résistance

Les bactéries multirésistantes les plus inquiétantes sont les entérobactéries multirésistantes – les entérobactéries comme *Escherichia coli* et *Klebsiella pneumoniae* sont des bactéries du tube digestif responsables d'un très grand nombre d'infections; *les staphylocoques dorés résistants à la méthicilline, les bacilles tuberculeux multirésistants, ou encore le bacille pyocyanique et les Acinetobacter baumanii* qui sont, des bactéries infectant les poumons de personnes atteintes de mucoviscidose et qui sont responsables d'infections nosocomiales (acquises en milieu de soin de santé, en particulier les hôpitaux et les cliniques).

La résistance aux antibiotiques ne concerne pas seulement les bactéries pathogènes. Toutes bactérie à l'état de compétence peut acquérir les gènes de résistances. De plus, les bactéries résistantes et les gènes de résistance peuvent se transmettre entre l'homme, les animaux et l'environnement. Ainsi, l'utilisation d'antibiotiques en médecine, vétérinaire et le rejet d'antibiotiques dans l'environnement contribuent à l'apparition de nouvelles souches bactériennes multirésistantes.



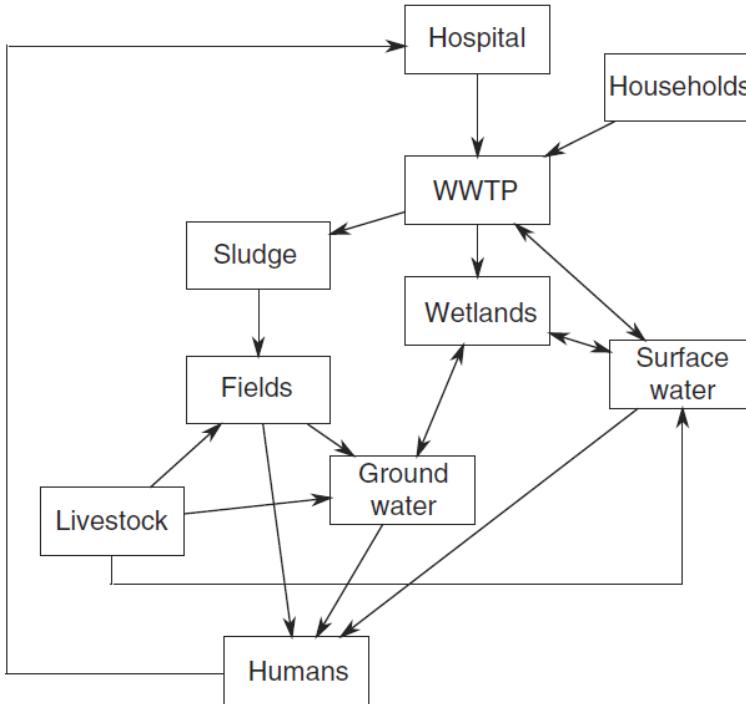
5.3. Dissémination des bactéries et gènes de résistance en milieux aquatique

The cause of transmission and the environmental proliferation of antibiotics and multi-resistant microbes are multiples and depend to the degree of the development of countries



➤ In developed countries

The technology of Waste Water Treatment Plant (WWTP) which receives the mixture of **effluent from hospitals** (urine and faeces from patients), **communities** (wastewater treatment system via people taking antibiotics from home), **animal farming** and **agricultural run-off**. The efficacy of WWTP can be considered a major point for antibiotics, antibiotic resistant bacteria (ARBs) and their genes (ARGs) for the aquatic environment



➤ In developing countries

The problem of ATB, ARB and ARG proliferation in non-clinical environment is considered as alarming for many reasons:

- No wastewater treatment
- The hospital effluents are discharged in receiving system without any previous treatment
- Open defecation
- No regulation of the prescription and use of antibiotics in human medicine and veterinary
- Use of contaminated water for irrigation and fresh produces

Congo DR



South of India

Impact of urban sewages in river receiving system : Trichy and Madurai (South India)

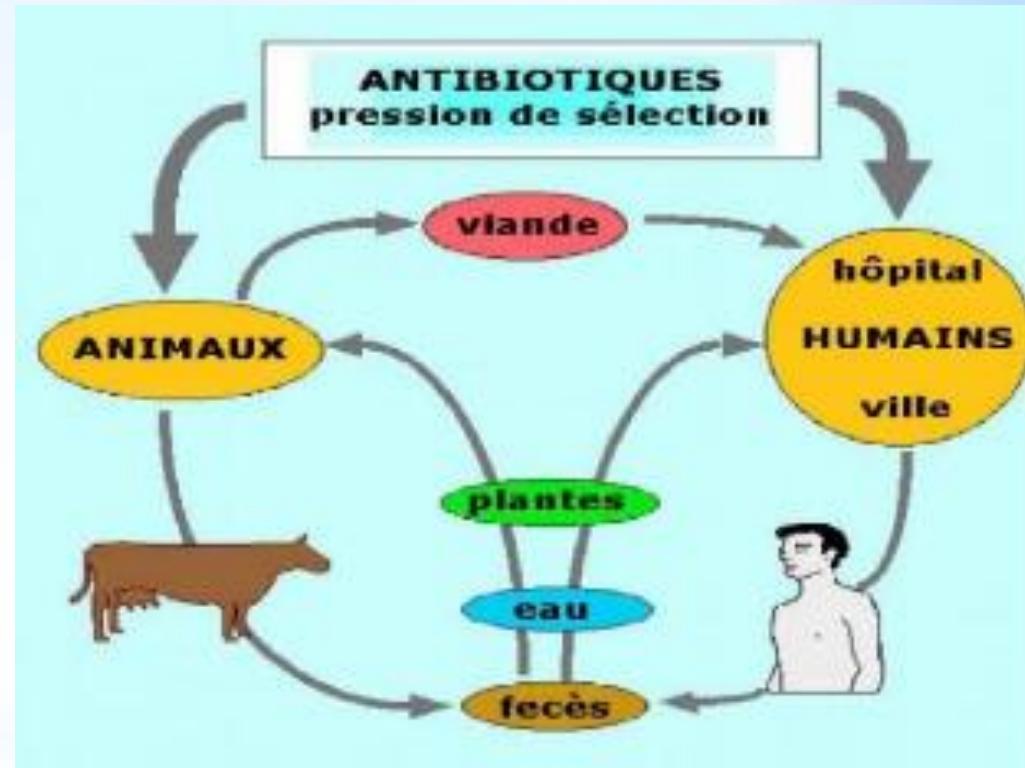
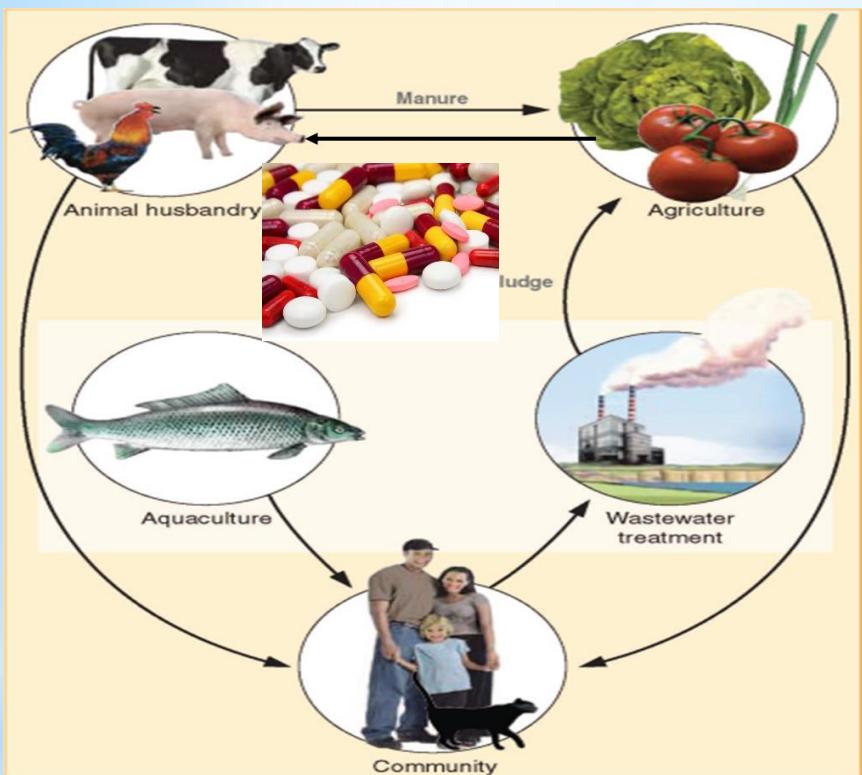


Taking antibiotics
for colds and flu?



*bla*_{NDM}

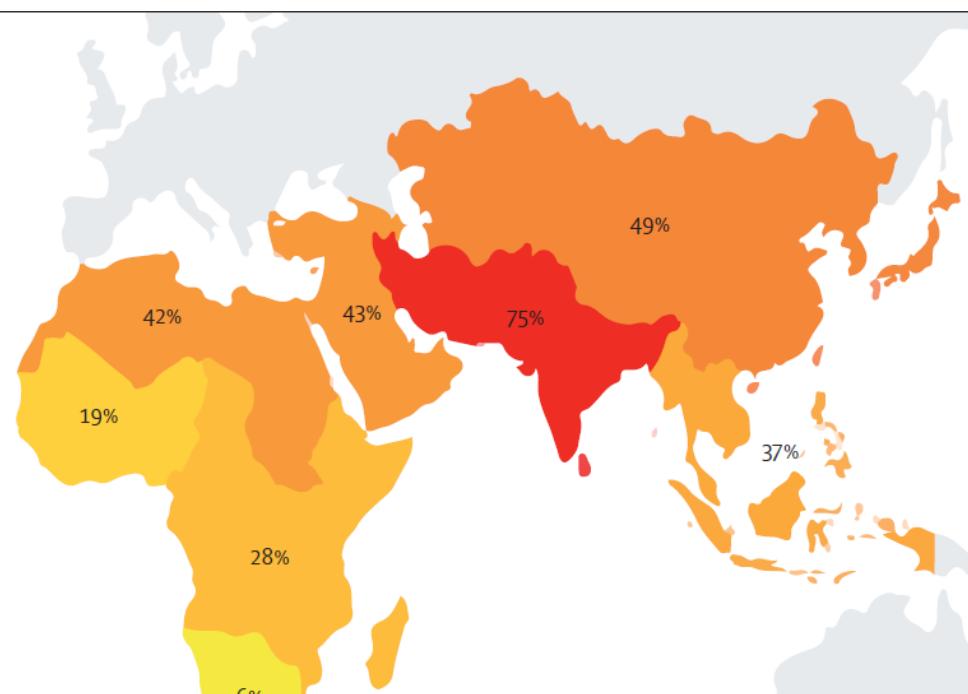
➤ Dissemination by contaminated and food habit



➤ Dissemination by travelers



Maris S Arcilla et al. *Lancet Infect Dis* 2017; 17: 78–85

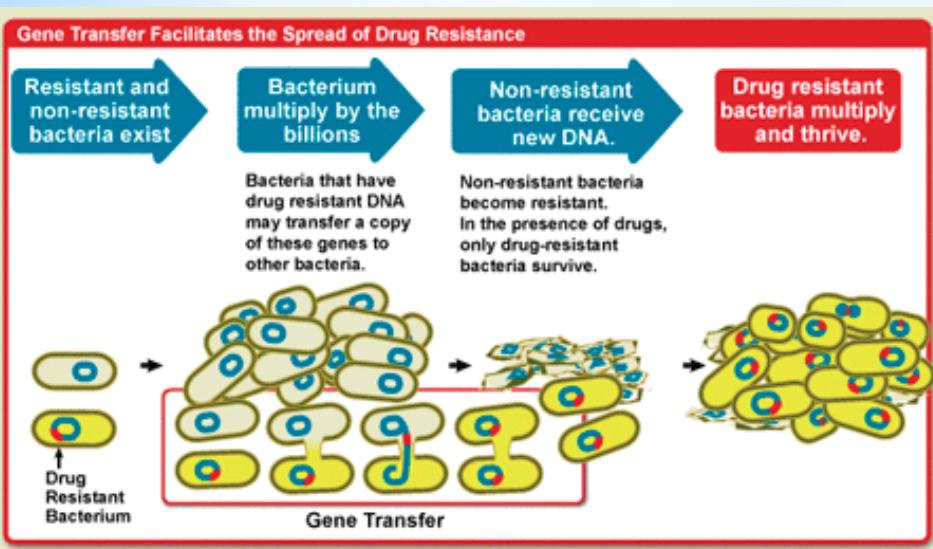


International travel contributes to the dissemination of antimicrobial resistance. Investigation of the acquisition of extended-spectrum β-lactamase-producing Enterobacteriaceae (ESBL-E) during international travel, with a focus on predictive factors for acquisition, duration of colonisation, and probability of onward transmission

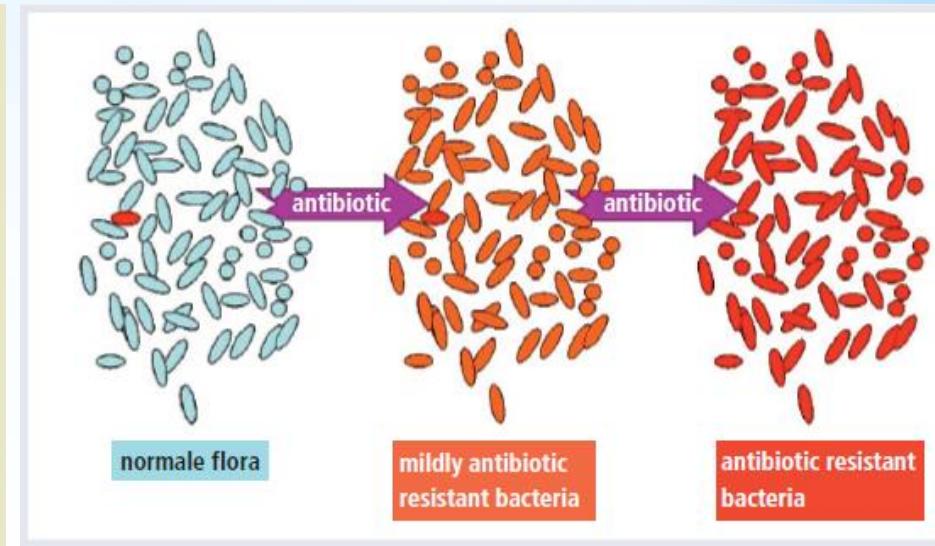
Figure 1: Percentages of travellers that acquired β-lactamase-producing Enterobacteriaceae per subregion, according to the United Nations geoscheme

5.4. Potential impacts

➤ Horizontal Gene Transfer (HGT) in environment

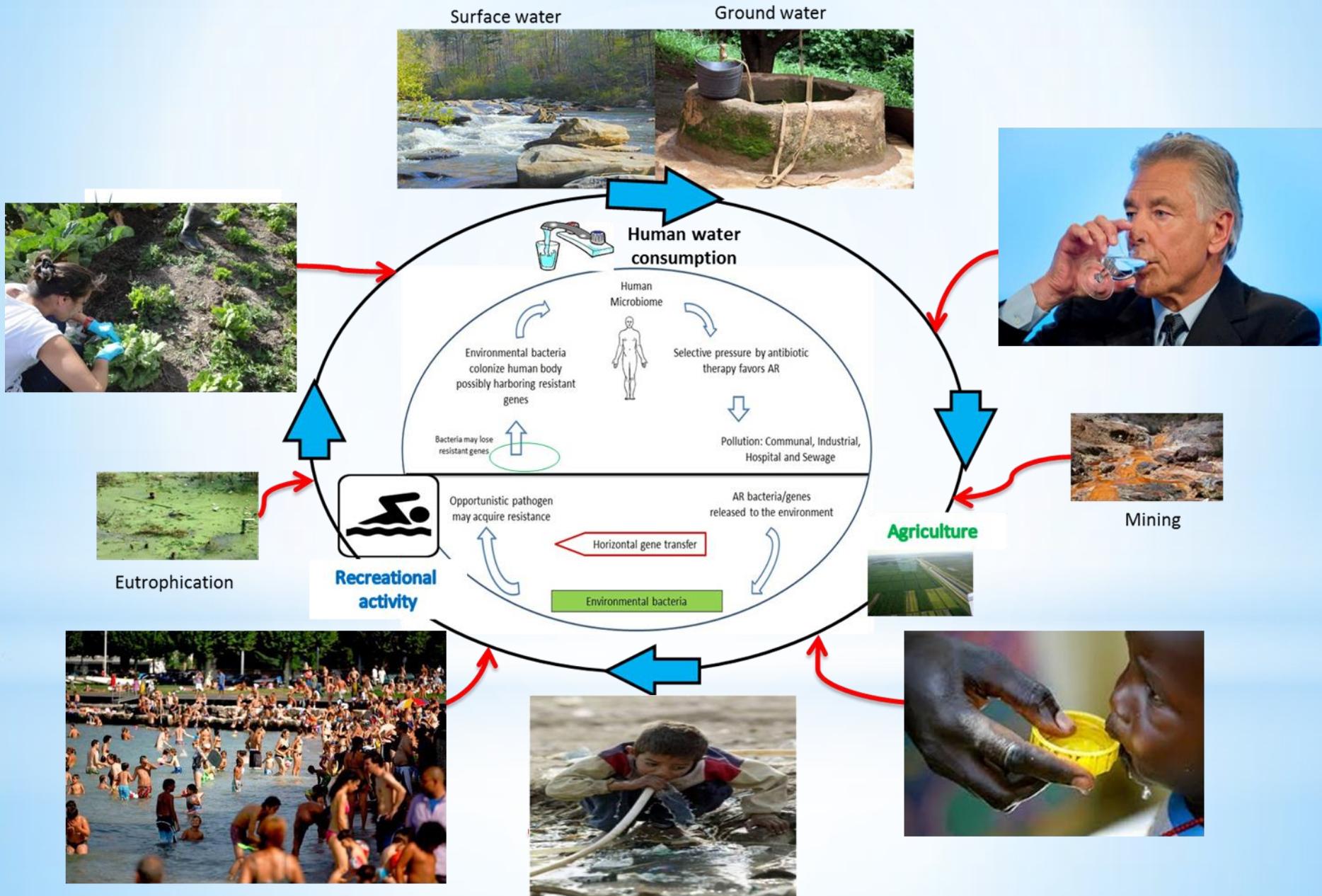


<https://www.quora.com/topic/Antibiotic-Resistance>



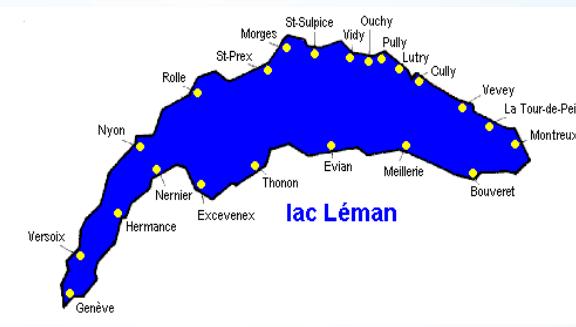
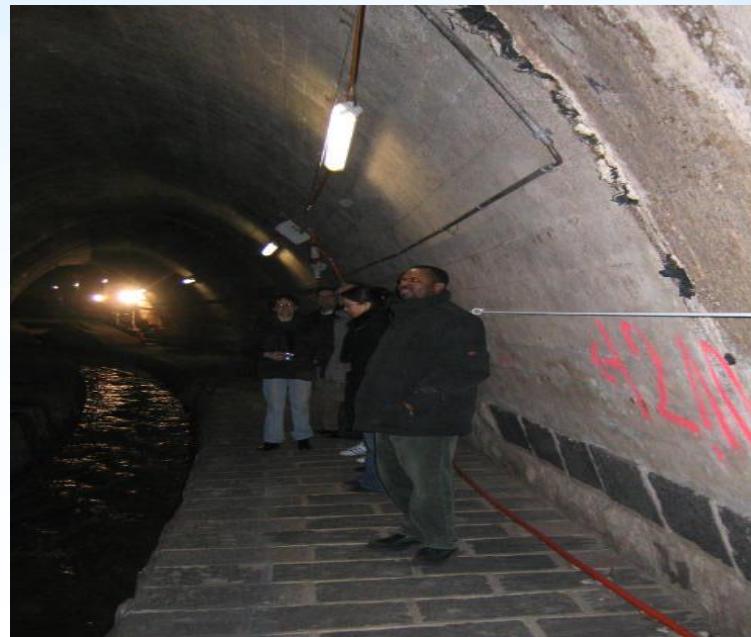
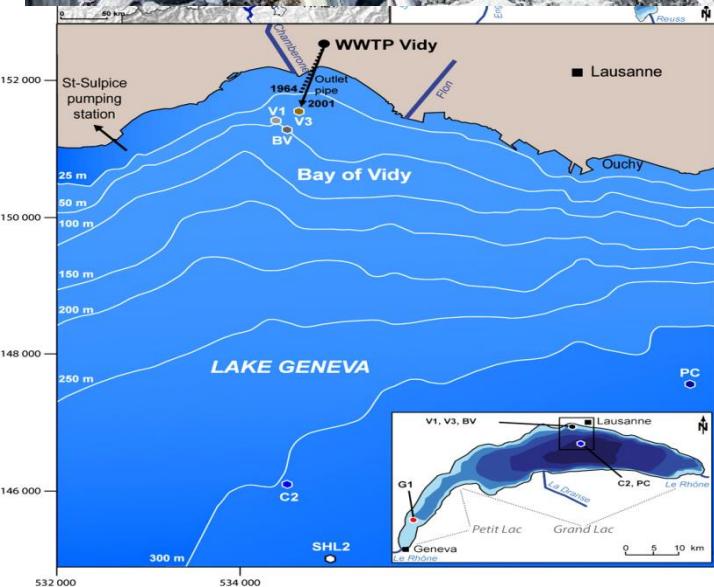
Horizontal gene transfer (HGT) between bacteria events are likely to be common in aquatic environments; integrons in particular are well suited for mediating environmental dissemination and proliferation of ARB and ARGs

➤ The return to the food and human chains

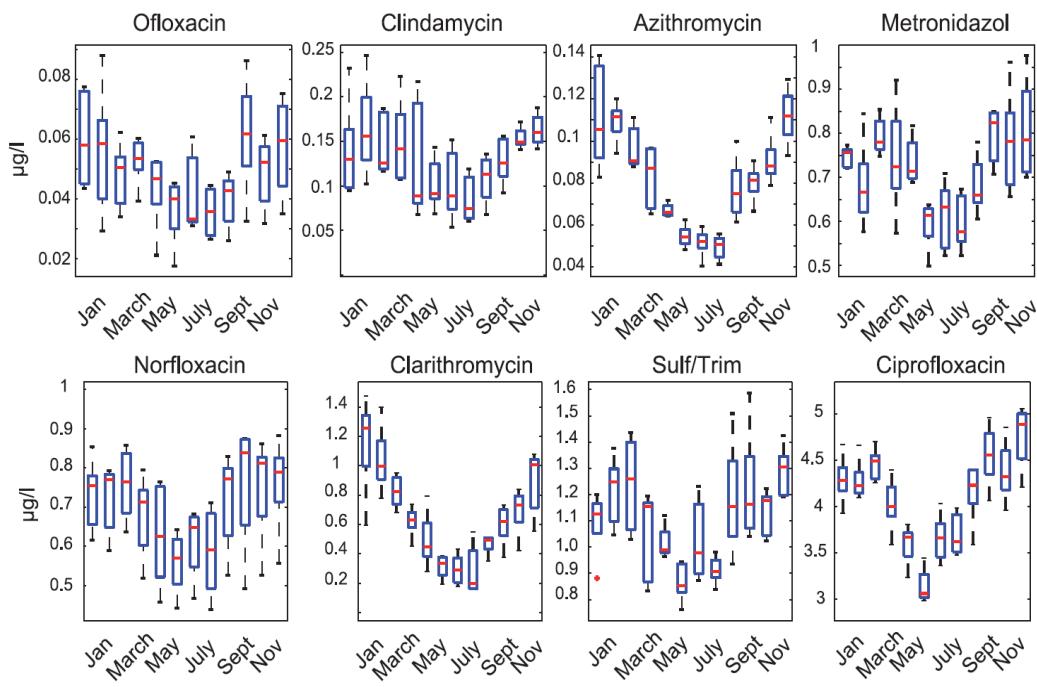
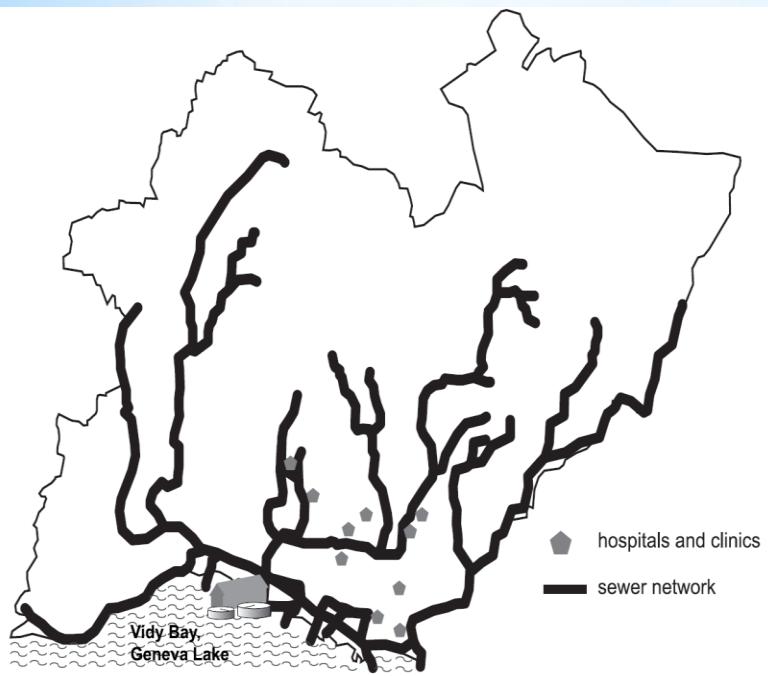


5.5. Case of Geneva Lake

➤ Vidy Bay Lausanne



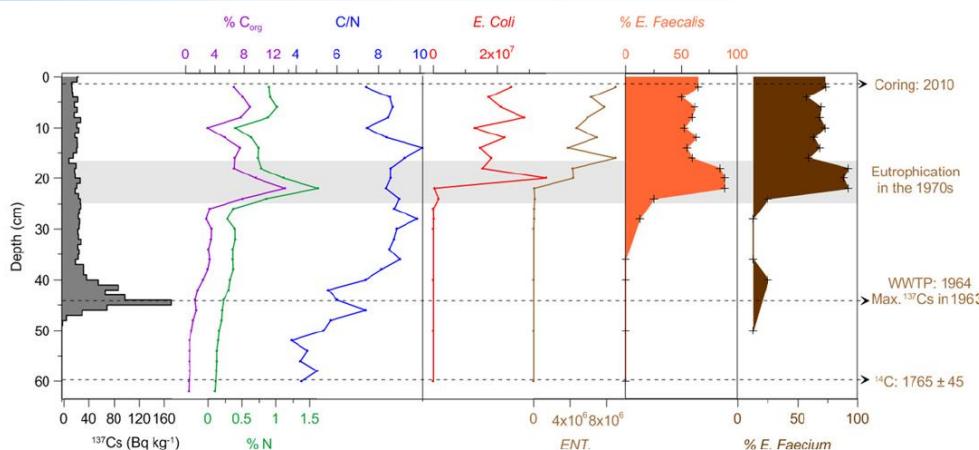
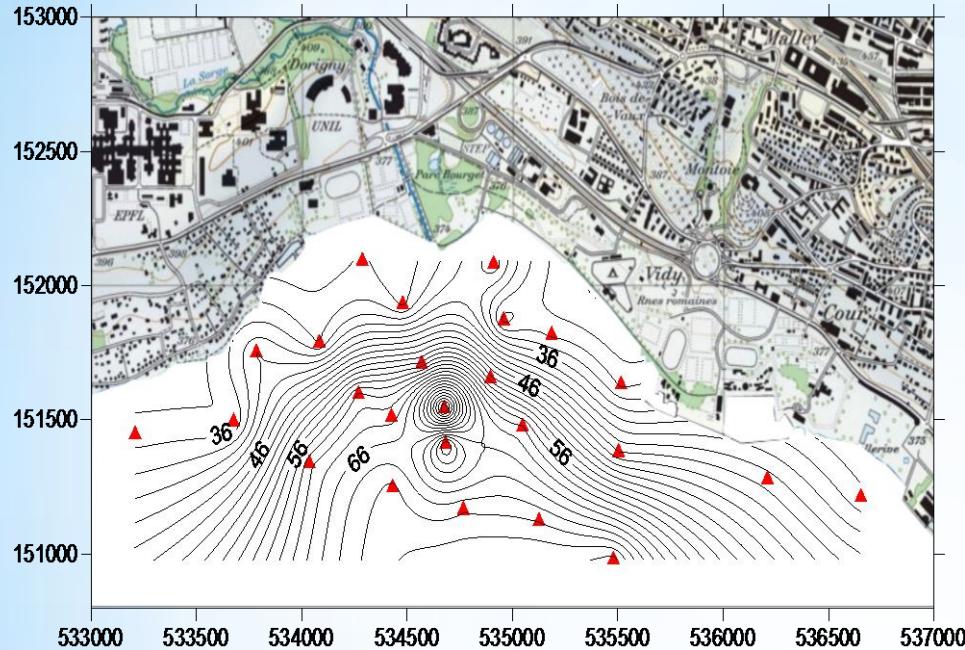
➤ Antibiotics Fluxes in Vidy Bay



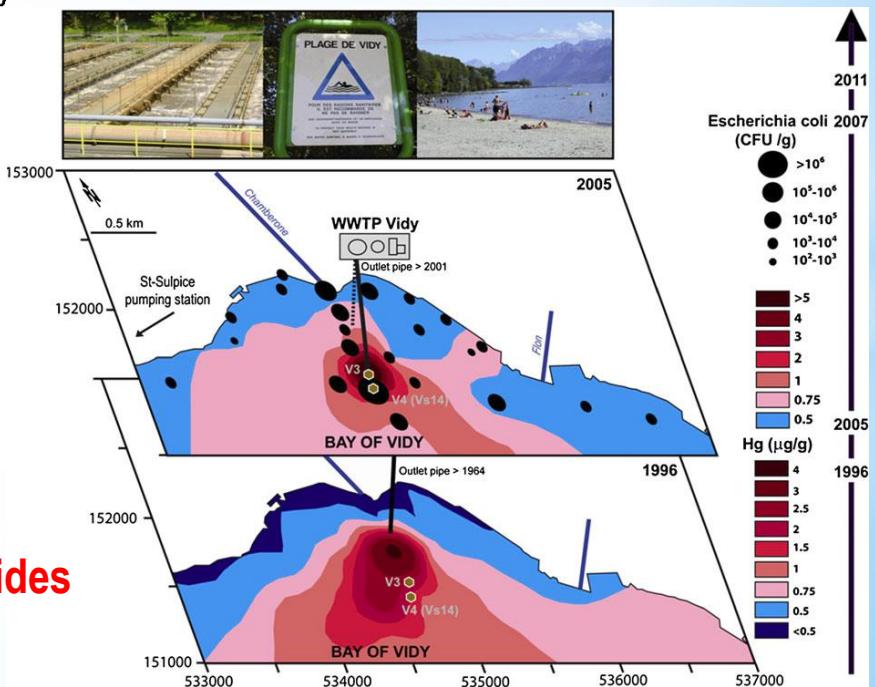
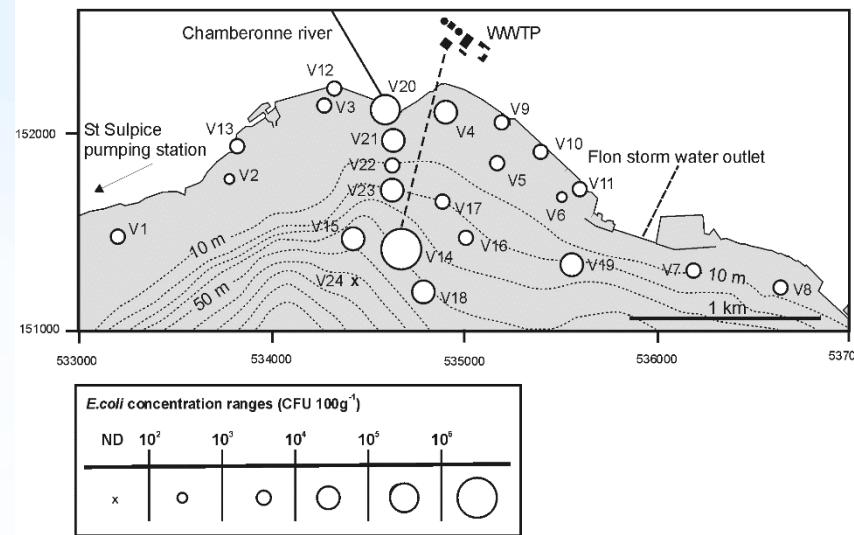
Investigation of seasonality of antibiotic concentrations in wastewater, with respect to WWTP network and hospital CHUV consumption . The study revealed important facts regarding antibiotic consumption as a source of environmental pollution.

Coutu S, Rossi L, Barry DA, Rudaz S, Vernaz N (2013). PLoS ONE 8(1): e53592.

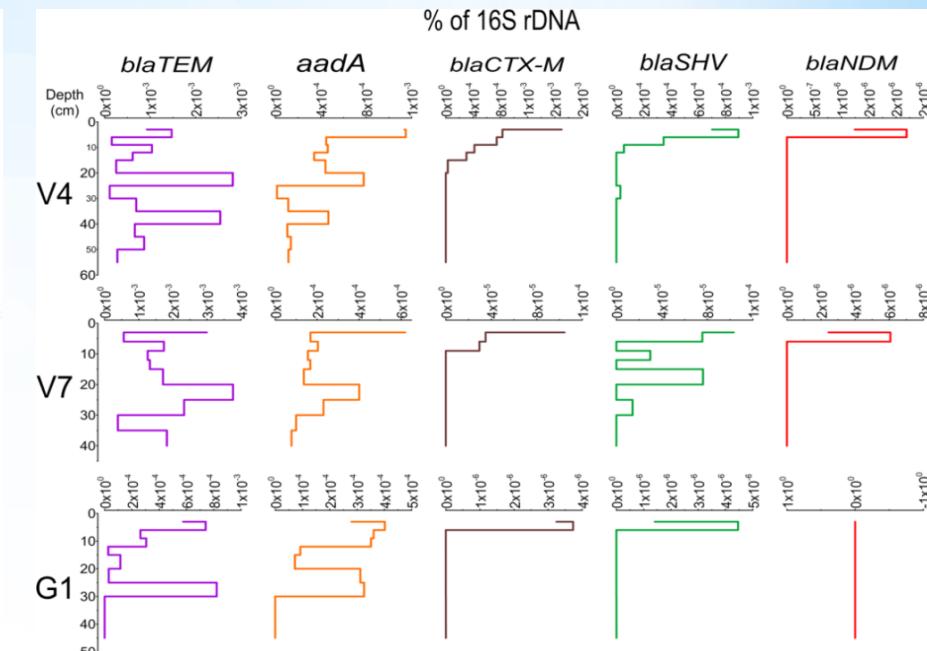
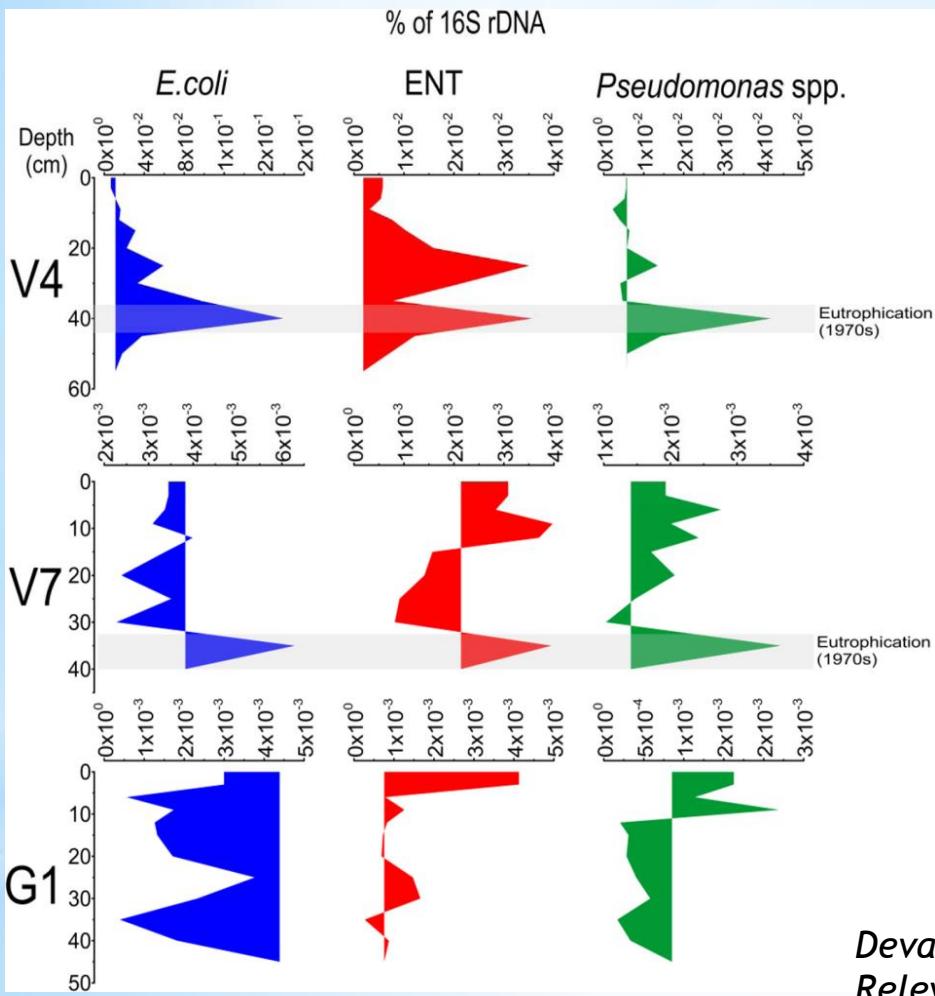
Pathogens distribution in water and surface sediments



92% of human bacteroides



➤ Clinically Bacteria and ARG resistant β -lactam genes dissemination

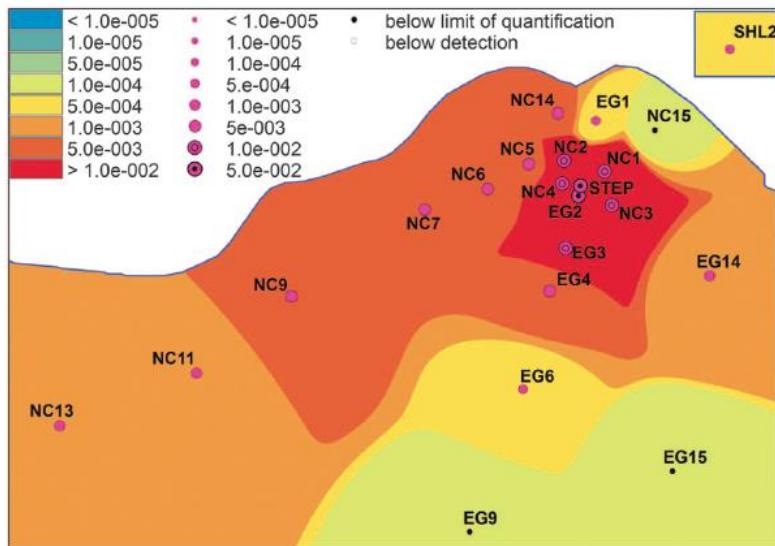


**Environmental
Science & Technology**

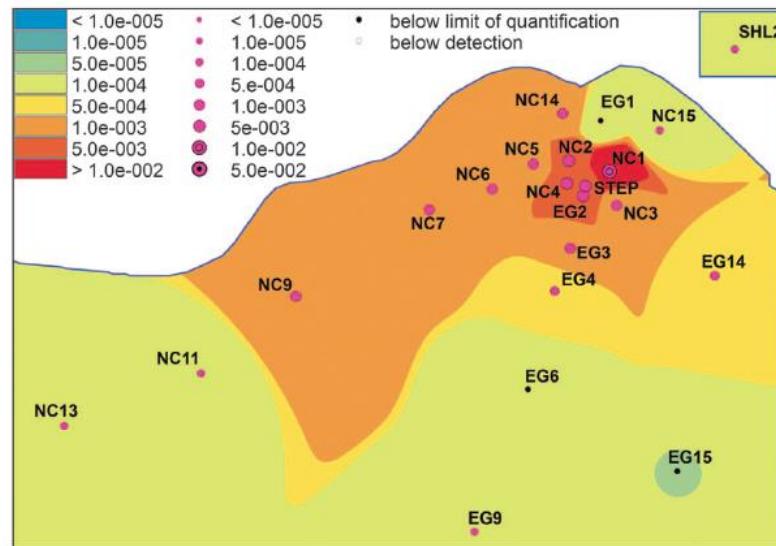
Devarajan,...,Poté, 2015. Accumulation of Clinically Relevant Antibiotic-Resistance Genes, Bacterial Load, and Metals in Freshwater Lake Sediments in Central Europe

➤ Other antibiotic genes dissemination in sediment of Vidy Bay

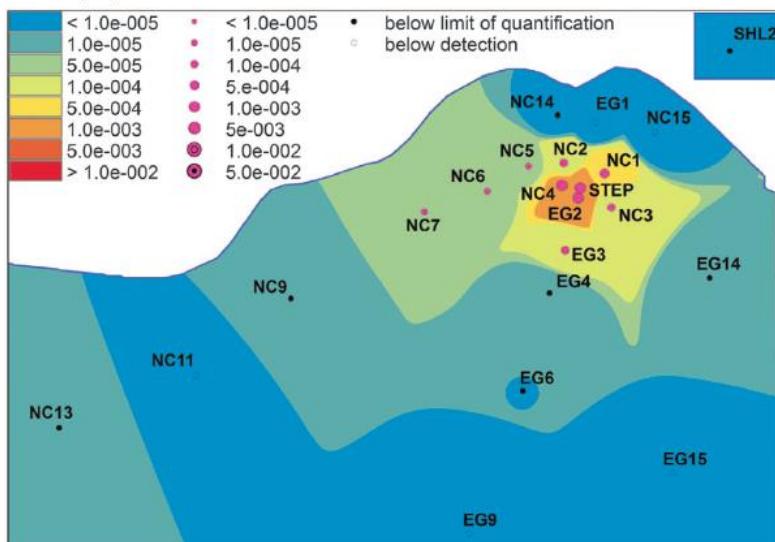
a *sul1*



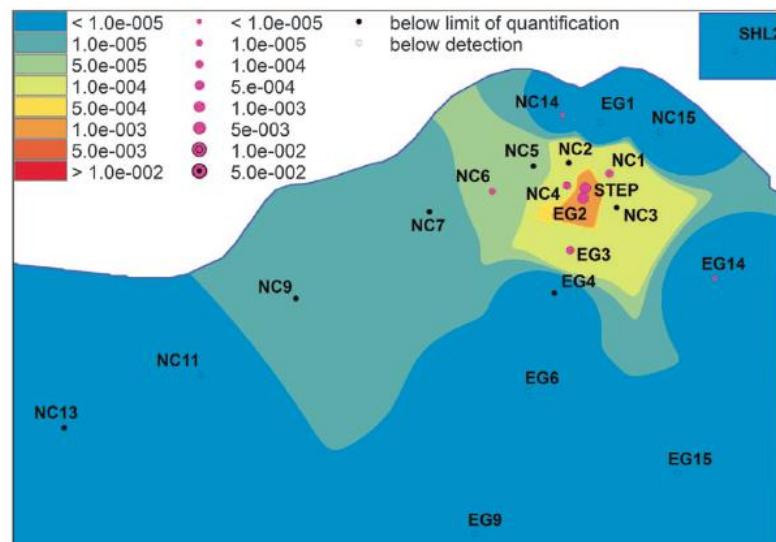
b *sul2*



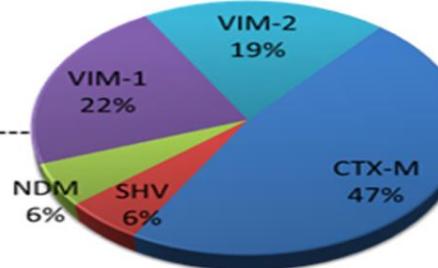
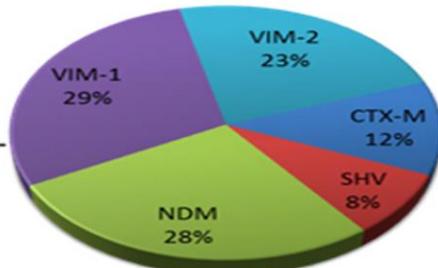
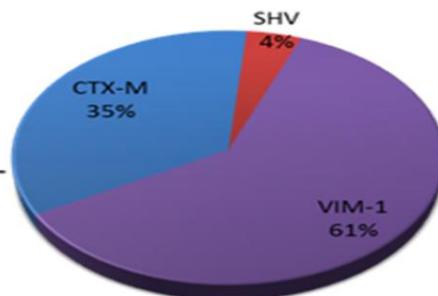
c *tet(W)*



d *tet(M)*



➤ Comparison Geneva and tropical conditions

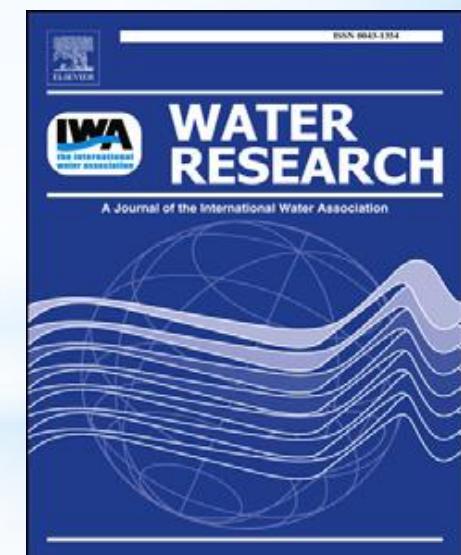


Antibiotic resistance against
Pseudomonas

India 35-60%

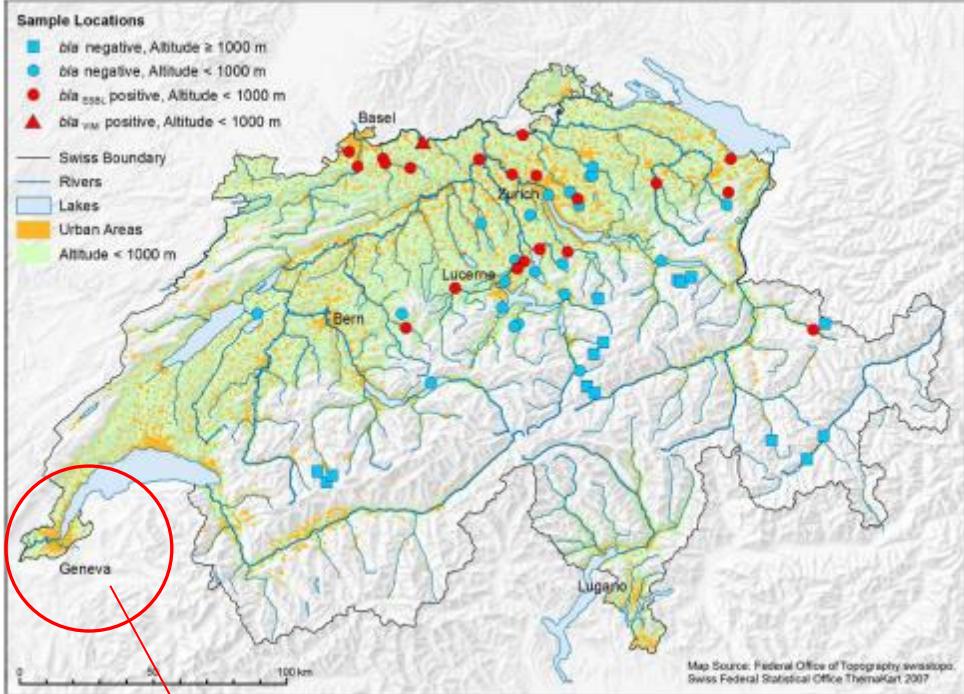
Congo 18-50%

CH 12-54%



Devarajan....Poté, 2017. Antibiotic resistant *Pseudomonas* spp. in the aquatic environment: A prevalence study under tropical and temperate climate conditions. Water Research 115 (2017) 256-265

5.6. Epidemiology of ESBL and CRE in Geneva Lake



| Bacterial species | Carbapenemase or 16S rRNA methylase | MWW | HW | TPI | TPE |
|-------------------------|-------------------------------------|-----|----|-----|-----|
| <i>E. coli</i> | OXA-48 | | | | |
| <i>E. coli</i> | OXA-181 | | | | |
| <i>E. coli</i> | OXA-48, VIM-1 | | | | |
| <i>E. coli</i> | NDM-5 | | | | |
| <i>E. coli</i> | ARMA | | | | |
| <i>E. coli</i> | RMTB | | | | |
| <i>K. pneumoniae</i> | OXA-48 | | | | |
| <i>K. pneumoniae</i> | NDM-9 | | | | |
| <i>K. pneumoniae</i> | KPC-2 | | | | |
| <i>Citrobacter</i> spp. | OXA-48 | | | | |
| <i>Citrobacter</i> spp. | OXA-48, NDM-1, ARMA | | | | |
| <i>Citrobacter</i> spp. | OXA-48, VIM-1 | | | | |
| <i>Citrobacter</i> spp. | VIM-1 | | | | |
| <i>E. cloacae</i> | OXA-48 | | | | |
| <i>E. aerogenes</i> | VIM-1 | | | | |

Zurfluh et al., 2017

No Quantitative data is available in Geneva canton

Zurfluh et al., 2013

Geneva, a cosmopolitan city with a foreign population of 40% provides a business friendly government and an outstanding quality of living.

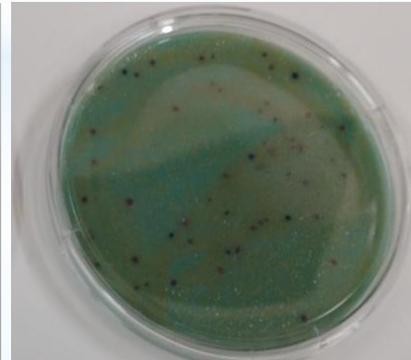
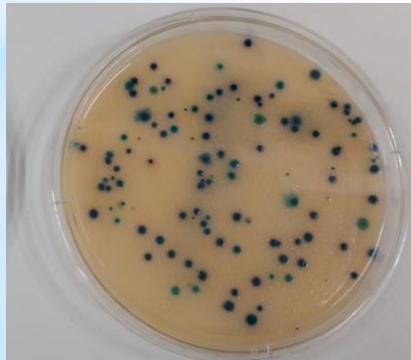
➤ Perspective study in Geneva beaches and Rhône



5.7. Potential Solution

There is very difficult to find appropriate solutions. Here we can give some hypothesis:

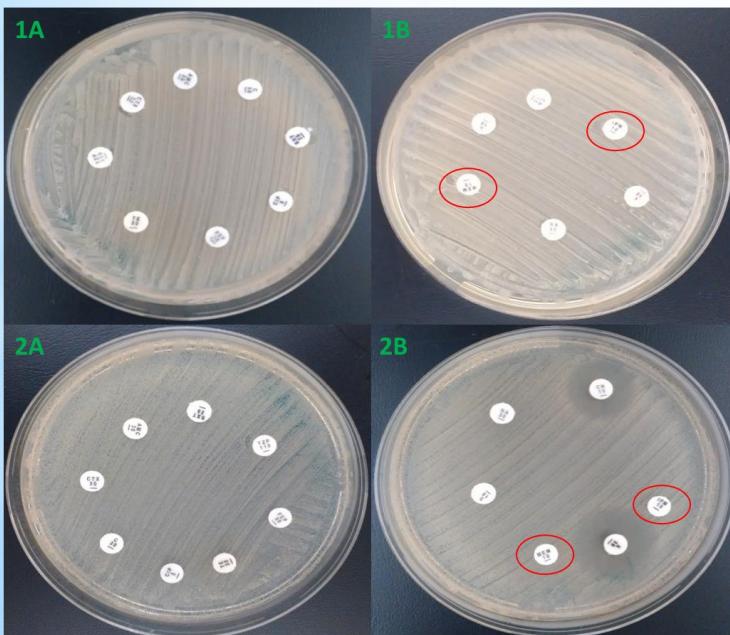
- Performing with research in different sittings
- Acquisition of quantitative data on the dissemination of antibiotic and antibiotic resistance according to the geographical location
- Regulation of the use of antibiotics in therapeutic: Clinical and veterinary
- Reduction of these biological contaminants and optimization of waste water treatment plants
- Education and sensibilisation in developing countries



**Bacterial isolates
FIB characterization
Susceptibility tests
Total DNA extraction
16S rRNA integrin class
Sequencing**

The epidemiology of antibiotic resistant in non-clinical environment is for these two last decades considered as worldly alarming problem. About 25000 persons die each year in EU and more than million in developing countries. The situation can be localized, but the effects and consequences are global !! We must act now !!!!

Financement



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DE LA RECHERCHE SCIENTIFIQUE



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Confédération suisse
Confederazione Svizzera
Confederaziun svizra

Office fédéral de l'environnement OFEV

THANK YOU