

### OBJECTIVES

- Identify key factors associated with the highest risk of virus emergence at the **4 stages of emergence**
- determine the **impact of the transmission route** on viral **evolutionary trajectories and cross-species transmission**
- unravel the mechanisms that govern **interactions** between a **virus, its hosts and the environment** to favour/limit cross-species transmission and adaptation to a new host
- evaluate most effective intervention strategies to **limit cross-species transmission and spread** in the new host
- **identify risk patterns of emergence** of practical relevance for disease surveillance, control and intervention, and pandemic preparedness

**Semi-flexibility clause** to include work on other viruses in area of expertise in response to emergence (MERS-CoV, Ebola Disease virus).

### Contacts

Sylvie van der WERF  
[sylvie.van-der-werf@pasteur.fr](mailto:sylvie.van-der-werf@pasteur.fr)

### PARTNERS

18 partners – 24 teams  
8 countries

1. Institut Pasteur (**coordinator**)
2. Istituto Zooprofilattico Sperimentale delle Venezie
3. Katholieke Universiteit Leuven
4. Aix Marseille Université
5. Eidgenössisches Departement des Innern
6. Imperial College of Science, Technology & Medicine
7. Agence Nationale de Sécurité Sanitaire de l'Alimentation, de l'Environnement et du Travail
8. Max Planck Gesellschaft zur Förderung der Wissenschaften e.V.
9. Universitat de Barcelona
10. Goeteborgs Universitet
11. Philipps Universität Marburg
12. Istituto Nazionale per le Malattie Infettive L. Spallanzani - IRCCS
13. The University of Edinburgh
14. Fondazione Istituto per l'Intercambio Scientifico
15. Foundation Health Sciences e-Training
16. Istituto Superiore di Sanità
17. Alma Mater Studiorum-Università di Bologna
18. Freunde von GISAID e.V.



### Lyssavirus

- What is the viral diversity in Europe and globally?
  - What ecologic, epidemiologic, virologic or host factors condition maintenance and spread of the virus in local ecosystems?
  - What factors determine cross-species transmission from bats to carnivores and humans?
- Ecological Factors Associated with European Bat Lyssavirus Seroprevalence in Spanish Bats and persistence mechanisms.  
*Serra-Cobo et al PLoS One. 2013*  
*J. Serra-Cobo UB & H. Bourhy IP*
- Rates of viral evolution are linked to host geography  
*Pons-Soler et al. PLoS ONE 2014*  
*P. Lemey KU Leuven*
- Introduction of fox rabies in Italy is due to 2 genetic groups  
*Fusaro et al IGE 2013*  
*G. Cattoli ISZvE*
- Mechanisms of innate immune response and immune evasion  
*Luco S et al PLoS Pathog. 2012*  
*Wiltzer et al. J Virol 2012*  
*Wiltzer et al. JID 2014*  
*H. Bourhy IP*

### Hepatitis E virus

- Viral diversity in Europe and globally
  - Ecologic, epidemiologic, virologic or host factors for:
    - maintenance and spread in local ecosystems
    - cross-species transmission between wild boars, pigs and humans (role of environment & food-chain)
- Novel hepatitis E like virus found in Swedish moose  
*Lin et al J Gen Virol 2014* *H. Norder UGOT*  
Hepatitis E Virus Genotype 4 Outbreak, Italy, 2011  
*Garbuglia et al EID 2013* *M.R. Capobianchi INMI*  
Hepatitis E Virus Genotype in a pig farm, Italy, 2013  
*Monne et al Epidemiol Infect 2014* *G. Cattoli ISZvE*  
Host factors modulated during HEV infection in swine  
*Rogee et al J. Virol. 2014* *N. Pavio ANSES*

### JEV & related (WNV, USUV, ZIKAV)

- Competence of European mosquitoes for JEV transmission ?
  - What animal could act as an amplifying host in Europe ?
  - What viral/host determinants determine the change of vector and host susceptibility ?
- Phylogeny of USUV from patient w/ neurological symptoms  
*Gaihani et al PLoS One 2013* *V. Sambridge UNIBO, E. Gould AMU*  
Novel flaviviruses from mosquitoes:mosquito-specific lineages  
*Huhtamo et al Virology 2014* *E. Gould, X. de Lamballerie AMU*  
RNA(+) viruses generated in days using infectious subgenomic replicons  
*Aubry et al JGV 2014* *E. Gould, X. de Lamballerie AMU*
- Sensitivity of WNV to the antiviral activity of IFNs  
*Bordj et al J Biol Regul & Hom Agents 2013*  
*M.R. Capobianchi INMI*

### CHIKV

- Extent and drivers of influenza virus diversity?
  - What factors determine potential for cross-species transmission and adaptation to humans?
  - What is the impact of the viral population (quasispecies) structure on cross-species transmission potential and pathogenicity?
- Structural determinants of IAV outside its host  
*Shigematsu et al. Influenza Other Respir Viruses, 2014*  
*Sawoo et al. PLoS ONE 2014* *M. Matrosovich UNIMAR*  
Switch from LP to HP for H7 viruses *Monne et al. JVI 2014* *G. Cattoli ISZvE*  
Synchronized global sweep of internal genes of modern avian influenza virus  
*Worobey et al. Nature 2014* *A. Rambaut UEDIN*
- Virus-host interactions using iPCA  
*Minier et al. Mol Cell Proteomics 2013*  
*Volker Czudai-Matwich et al. JVI 2014* *M. Matrosovich UNIMAR*  
Avian influenza A virus PB2 promotes IFN induction of sIV in swDC  
*Ocana-Macchi et al. Virology 2012* *A. Summerfield FDEA-IVI*  
Cellular and humoral cross-immunity against H3N2 in HIV + subjects  
*Agati et al. PLoS ONE 2014* *M.R. Capobianchi INMI*  
Prediction of global transmission of influenza H3N2  
*Lemey et al. PLoS Pathogens 2014* *P. Lemey KU-Leuven*

### Influenza A virus

### EBOV

- Mobile labs and diagnostics
  - Virus evolution and origin
  - Host-virus interactions
  - Transmission dynamics & parameters
- Contribution to mobile labs & diagnostics *J.-C. Manuguerra IP; G. Ipolito INMI*  
Genomic surveillance elucidates Ebola virus origin and transmission during the 2014 outbreak *Gire et al. Science 2014* *A. Rambaut UEDIN*  
Phylogenetic analysis of Guinea 2014 EBOV outbreak  
*Dudas et al. PLoS Current 2014* *A. Rambaut UEDIN*
- Ebola Virus Disease in West Africa – The first nine months of the epidemic  
*WHO Ebola Response Team NEJM 2014*  
West African Ebola Epidemic after One Year-  
*WHO Ebola Response Team N Engl J Med 2014*  
Impact of travel restrictions on international Ebola spread  
*Poletto et al. Eurosurveillance 2014* *V. Colizza ISI*

### MERS-CoV

- Clinical features
- sites and kinetics of virus shedding
- Virus evolution and origin
- Parameters of transmission

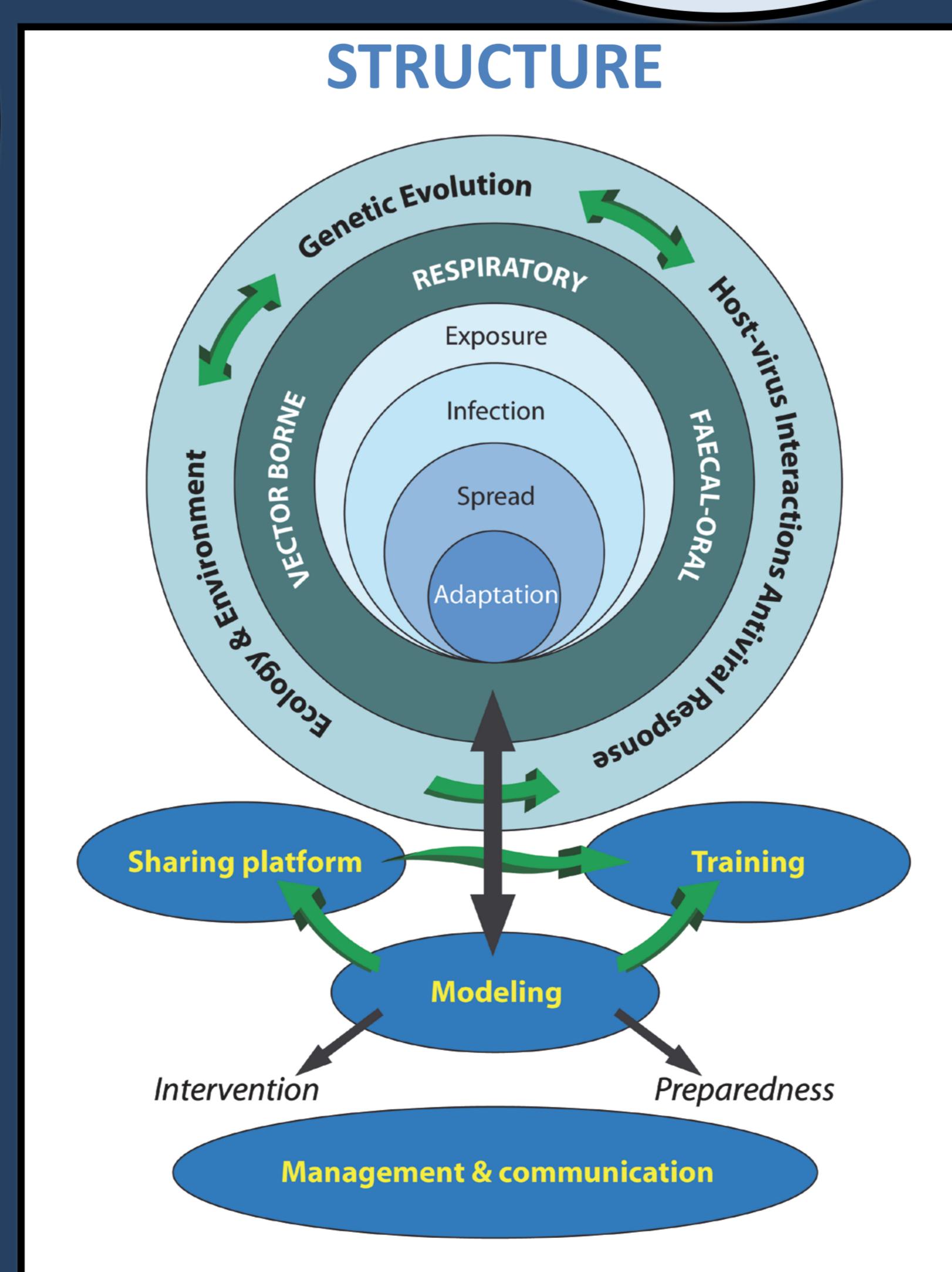
- Clinical features and viral diagnosis of two cases of infection with Middle East Respiratory Syndrome coronavirus:  
a report of nosocomial transmission.  
*Gury et al. Lancet. 2013*  
Transmission and evolution of the Middle East respiratory syndrome coronavirus in Saudi Arabia: a descriptive genomic study.  
*Cotten et al. Lancet. 2013* *A. Rambaut UEDIN*
- Transmission scenarios for Middle East Respiratory Syndrome Coronavirus (MERS-CoV) and how to tell them apart.  
*Breban et al. Lancet. 2013*  
Interhuman transmissibility of Middle East respiratory syndrome coronavirus: estimation of pandemic risk.  
*C. Donnelly IMPERIAL S. Cauchemez IP*

### Modelling

- Mapping methods for animal densities & behaviours
- Transmission models
- Models for risk assessment and contingency planning
- Evolutionary epidemic models

- Rabies and CDV epidemics in rd fox population in Italy  
*Nouvellet et al PLoS One 2013* *C. Donnelly IMPERIAL*  
Role of environmental compartment for IAV transmission  
*R. Breban IP*  
*Breban J Math Biol 2012*
- Surveillance to estimate epidemic potential of emerging zoonoses : H3N2  
*Cauchemez et al. PLoS Med 2013* *C. Donnelly IMPERIAL S. Cauchemez IP*
- Host mobility drives pathogen competition in spatially structured populations  
*Poletto et al PLoS Comput Biol 2013* *V. Colizza ISI*  
Exposure and human-to-human transmission for emerging pathogens  
*Kucharski et al PLoS Comput Outbreaks 2014* *V. Colizza ISI*  
Local and regional spread of chikungunya fever in the Americas  
*Cauchemez et al Eurosurveillance 2014* *S. Cauchemez IP*

### STRUCTURE



### TRAINING

#### Scientist exchange Program

#### Training courses and workshops

-Zoonotic features of IAV, HEV, JEV, LYS



-Virus evolution and phylogeny

-Rabies hands-on course (WHO, IP Int network)

-GISAID-WHO Training workshop : Geneticx analyses of influenza viruses"

#### E-learning

### DISSEMINATION

- Interactions with other EU projects (ANTIGONE, PREPARE, SILVER, FLUPHARM, ...)
- Outreach to main stake-holders (WHO, OIE, ECDC,..)