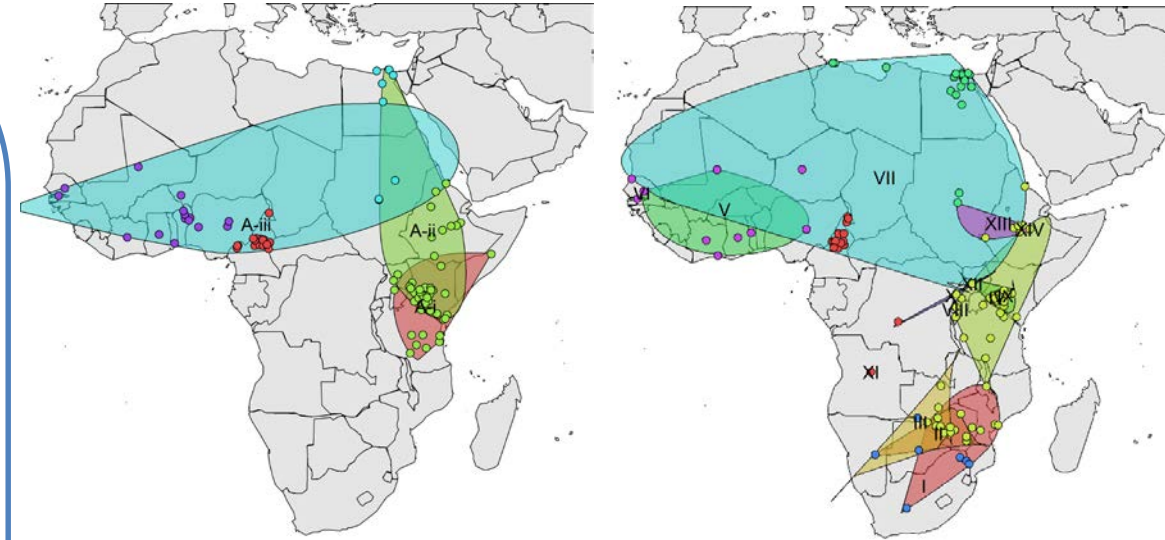


# A NOVEL MODELLING APPROACH FOR ENDEMIC FMD IN SUB-SAHARAN AFRICA

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### FMD subtypes and topotypes in Africa

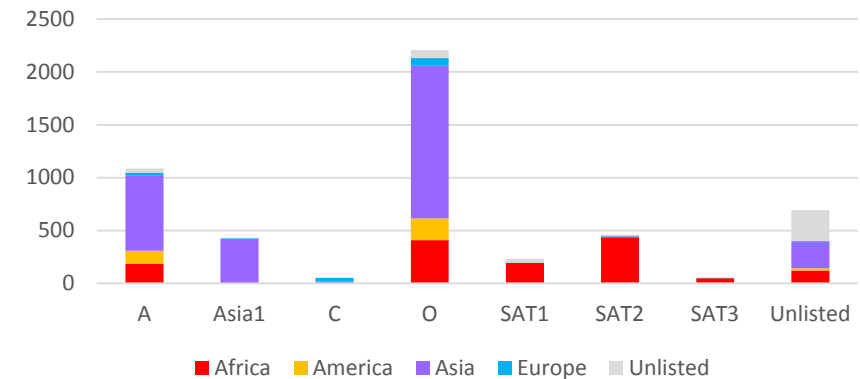
- Control of foot-and-mouth disease  
→ understand the epidemiology of the disease
- Interested in :
  - **Transmission** rates between **locations**
  - Number of transition between locations
  - **Effect** of anthropological/ environmental **variable**
- Overlap in FMDV serotypes and topotypes in **West, Central and East Africa**
  - Serotypes O, A, SAT 1, SAT 2
  - Great circulation between countries
- Epidemiology of FMD in Africa **still unclear**
  - Role of livestock and wildlife?
  - Lack of data



*FMDV A clades distribution in Africa*

*FMDV SAT2 topotypes distribution in Africa*

### Number of VP1 (1D) Sequences in Genbank

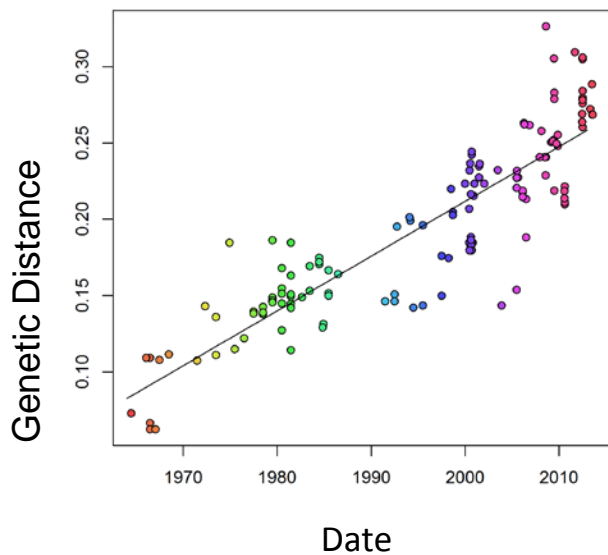


Pathogen sequences accumulate mutations over time

Tree of FMD A



Genetic Distance from Root



Evolutionary tree timescale by a molecular clock

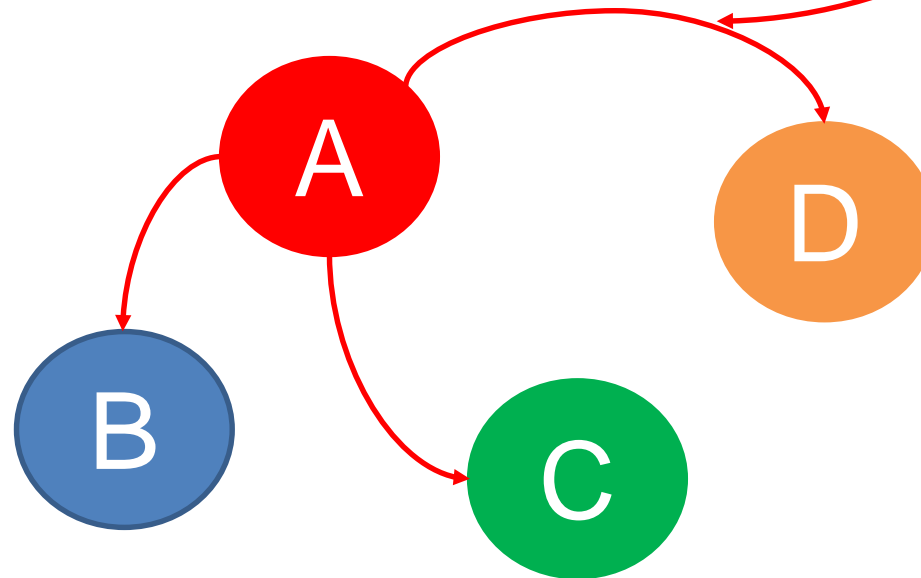
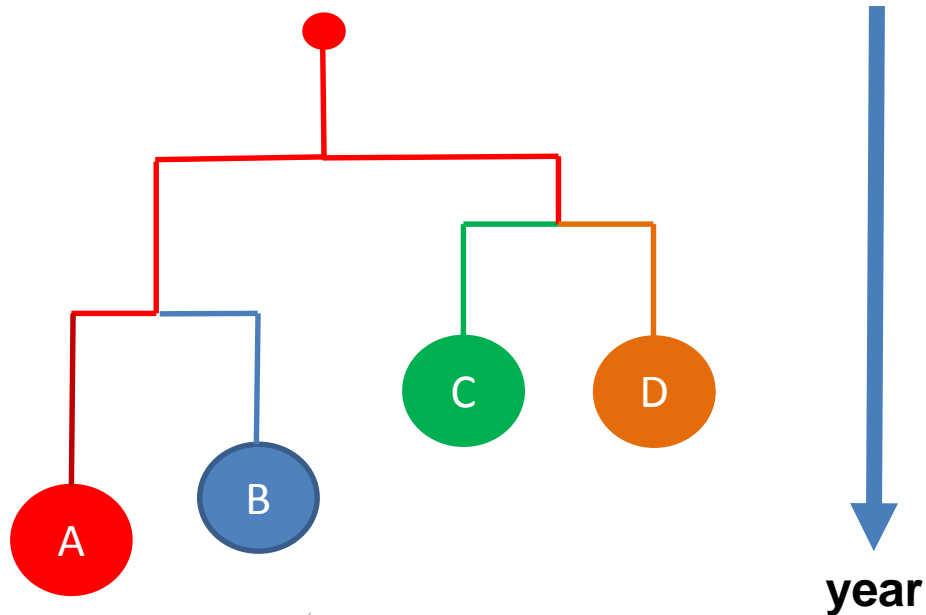
Time to most recent common ancestor



## Integration of spatial data

- Add **locations** to phylogenetic tree → evolutionary trait
- Estimate transition rates between **locations** along branches
- Transmission pattern between **locations** represented by rate matrix

|   | A              | B              | C              | D              |
|---|----------------|----------------|----------------|----------------|
| A | 0              | $\lambda_{ab}$ | $\lambda_{ac}$ | $\lambda_{ad}$ |
| B | $\lambda_{ba}$ | 0              | $\lambda_{bc}$ | $\lambda_{bd}$ |
| C | $\lambda_{ca}$ | $\lambda_{cb}$ | 0              | $\lambda_{cd}$ |
| D | $\lambda_{da}$ | $\lambda_{db}$ | $\lambda_{dc}$ | 0              |



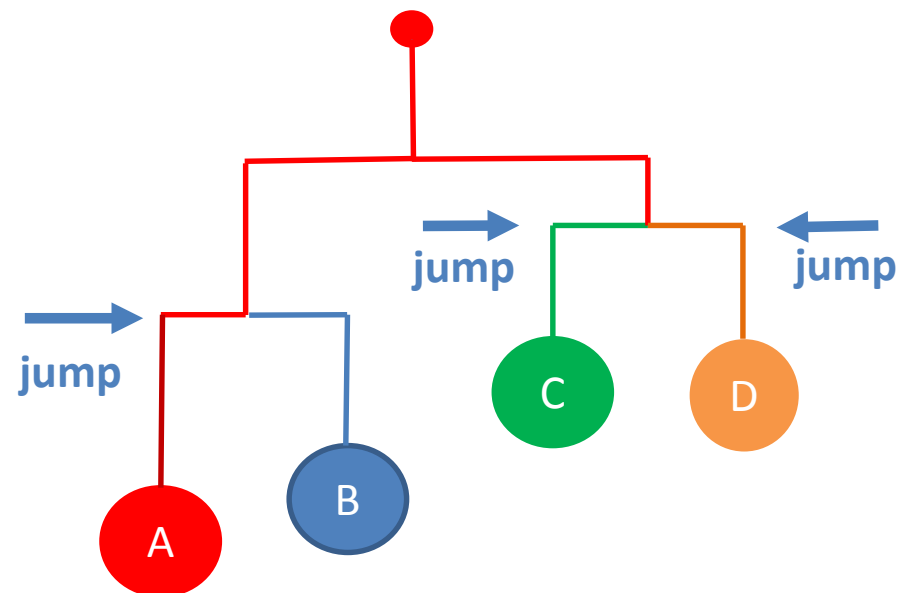
**BSSVS ( Bayesian Stochastic Search Variable Selection )**

- Determine well supported location transmission rate
- Many location transitions are unlikely → reduce the number of rates
  - Model augmented with indicator variables  $\Delta_{ijk}$ .  
 $\Delta_{ijk} = 0 \rightarrow$  rate between states  $j$  and  $k$  is 0.  
 $\Delta_{ijk} = 1 \rightarrow$  rate between states  $j$  and  $k$  is  $\lambda_{ijk}$
- Provide a **Bayes Factor** → tests for well supported individual rates

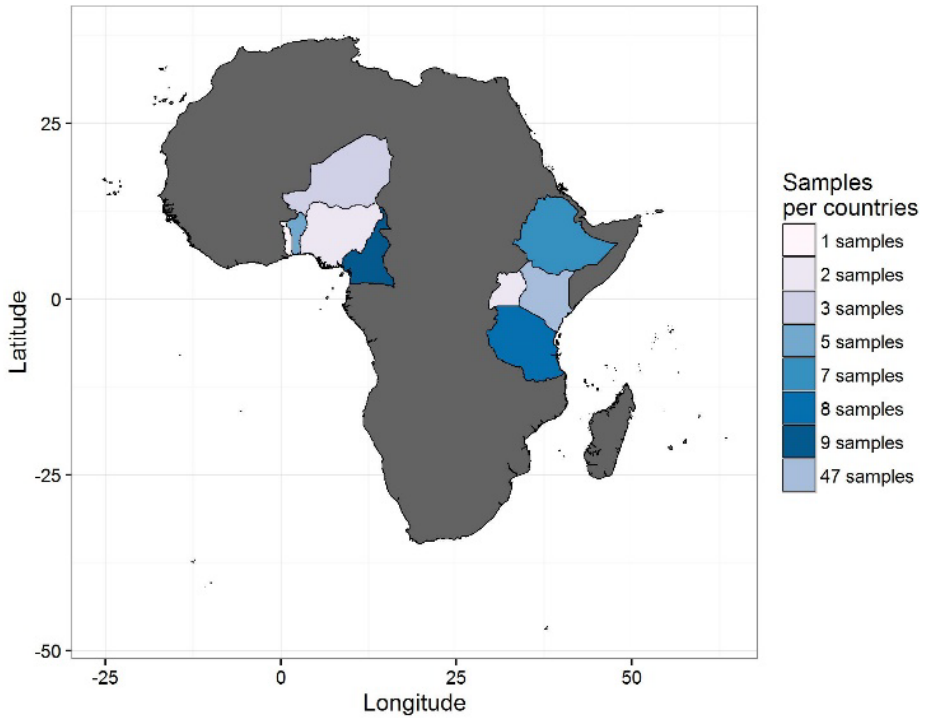
**Markov jump**

- Estimate the number of **location** transitions (Markov jumps) along the branches of the unknown tree

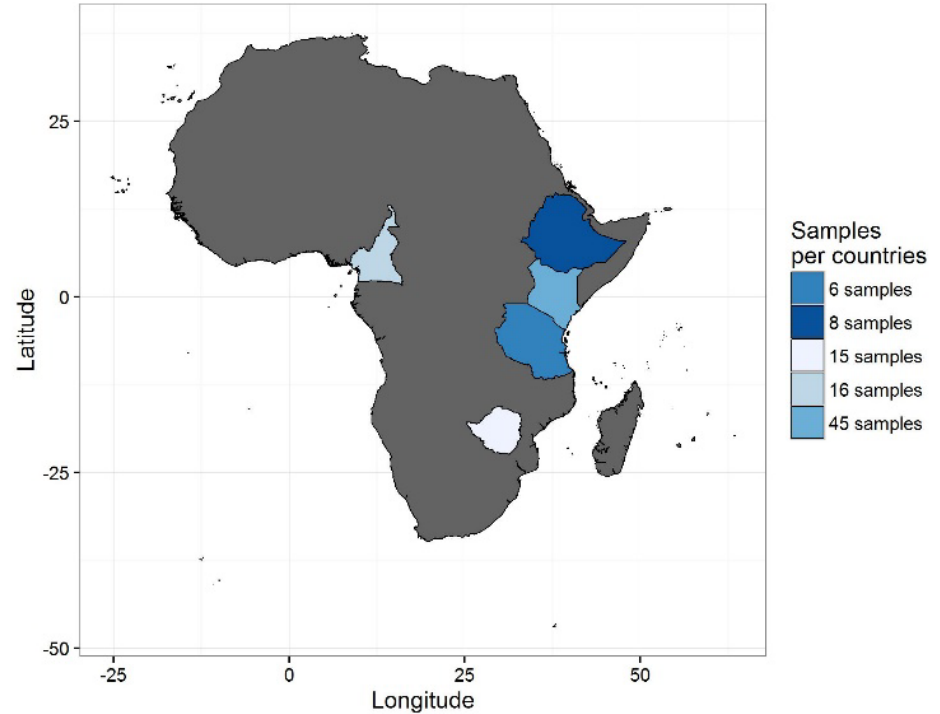
|   | A                          | B                          | C                          | D                          |
|---|----------------------------|----------------------------|----------------------------|----------------------------|
| A | 0                          | $\Delta_{ab} \lambda_{ab}$ | $\Delta_{ac} \lambda_{ac}$ | $\Delta_{ad} \lambda_{ad}$ |
| B | $\Delta_{ba} \lambda_{ba}$ | 0                          | $\Delta_{bc} \lambda_{bc}$ | $\Delta_{bd} \lambda_{bd}$ |
| C | $\Delta_{ca} \lambda_{ca}$ | $\Delta_{cb} \lambda_{cb}$ | 0                          | $\Delta_{cd} \lambda_{cd}$ |
| D | $\Delta_{da} \lambda_{da}$ | $\Delta_{db} \lambda_{db}$ | $\Delta_{dc} \lambda_{dc}$ | 0                          |



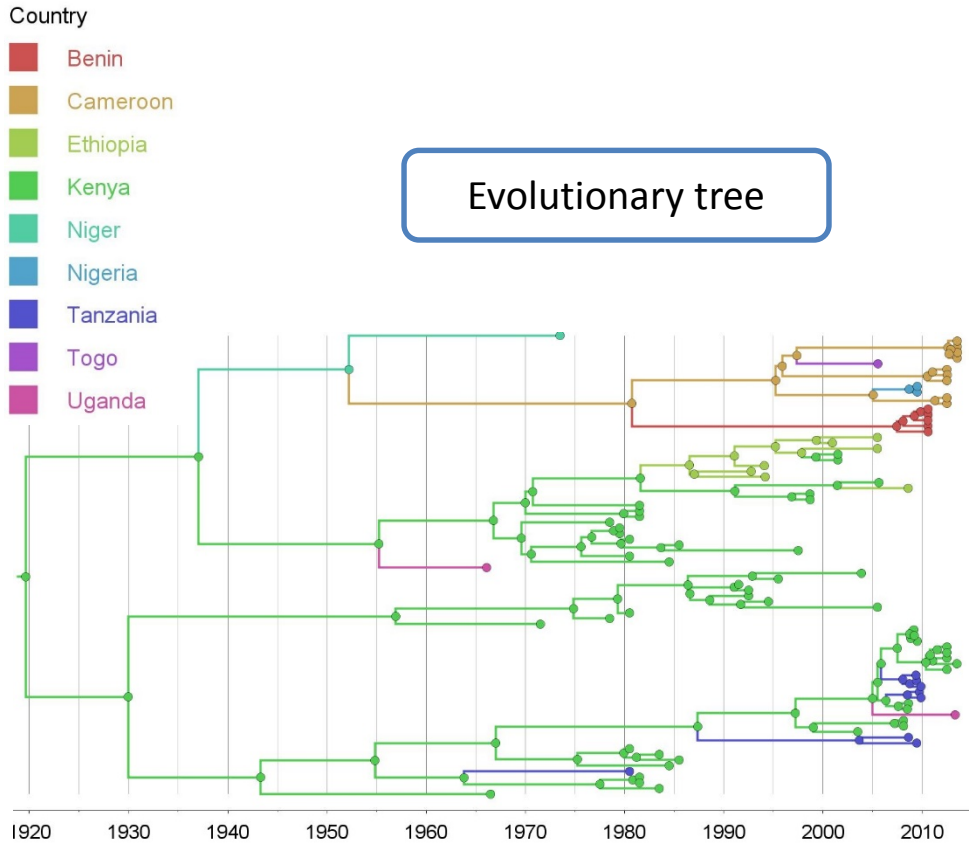
FMDV A



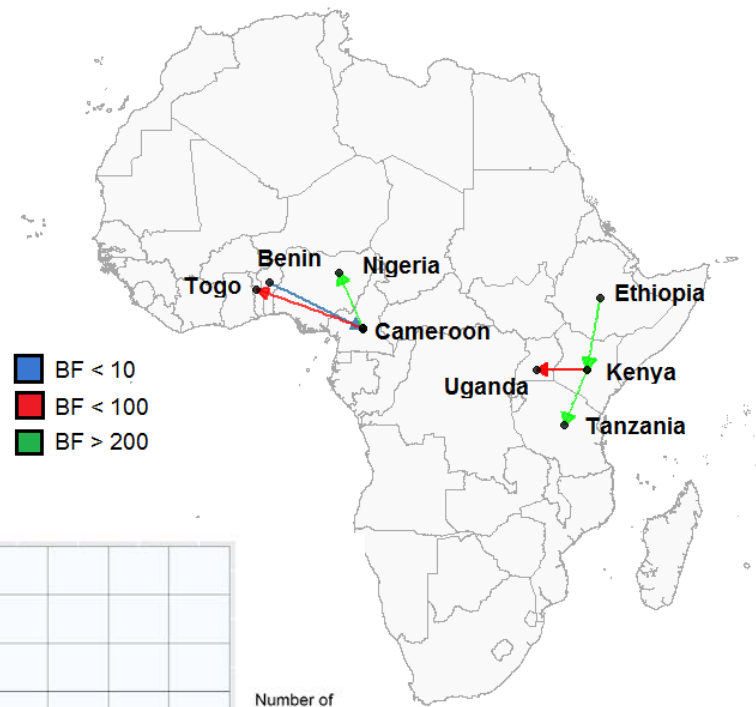
FMDV SAT2



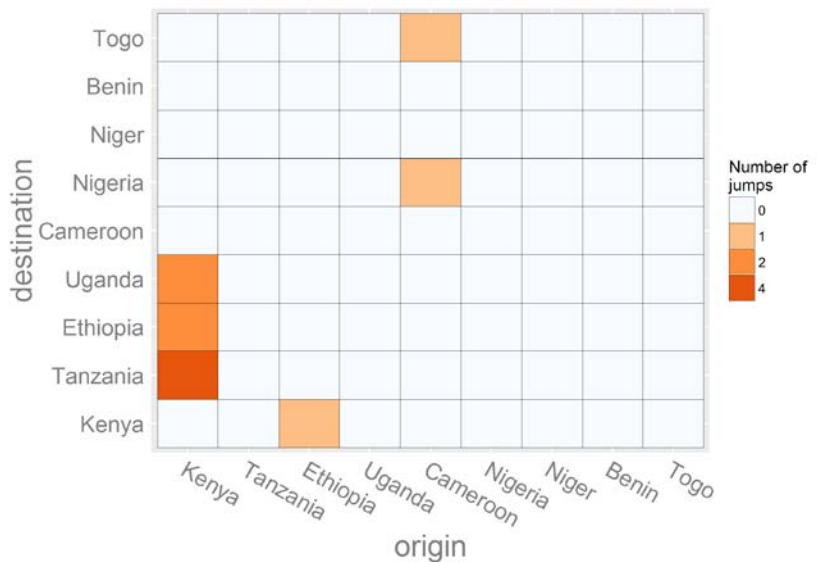
## FMDV A



## Supported routes of transmission



## Markov jumps



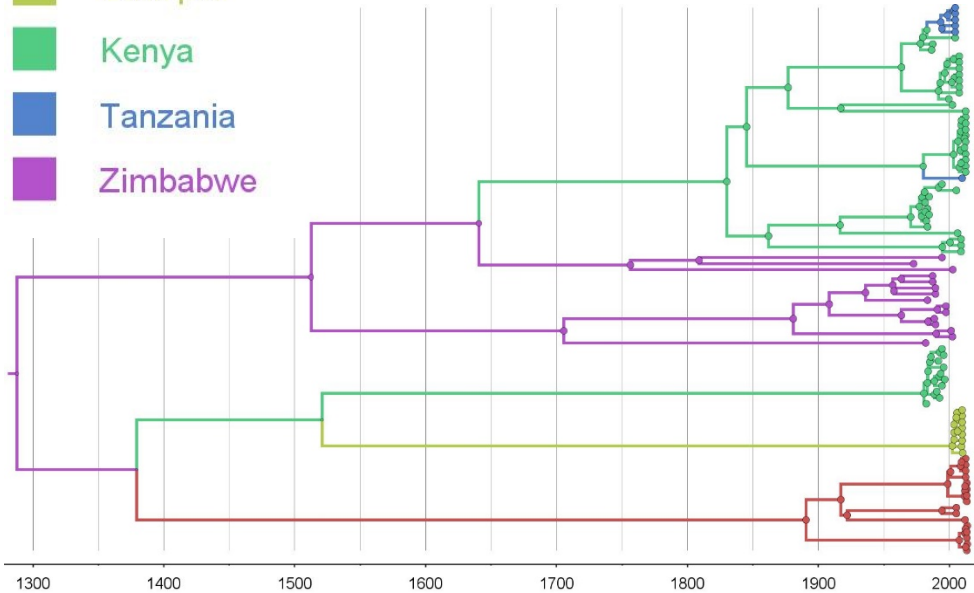
FMDV SAT2

Supported routes of transmission

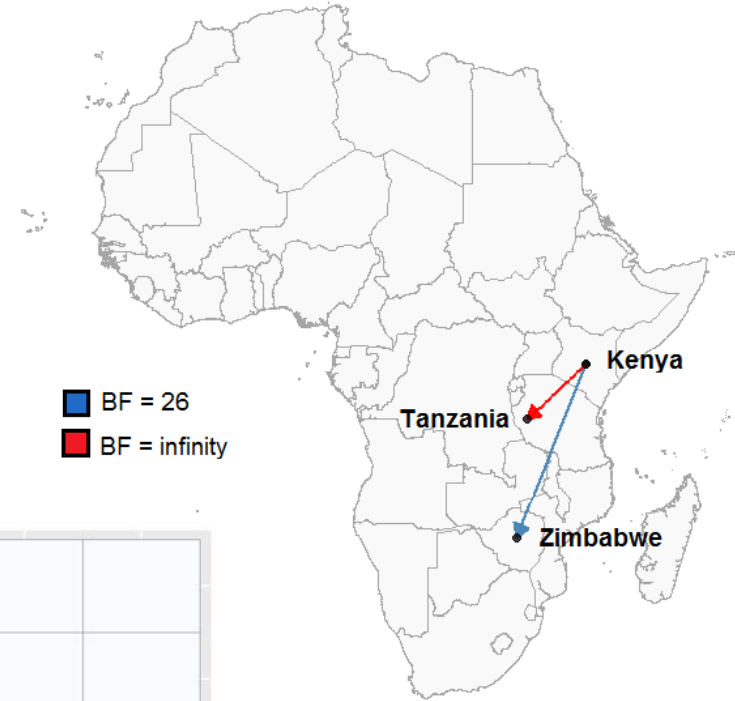
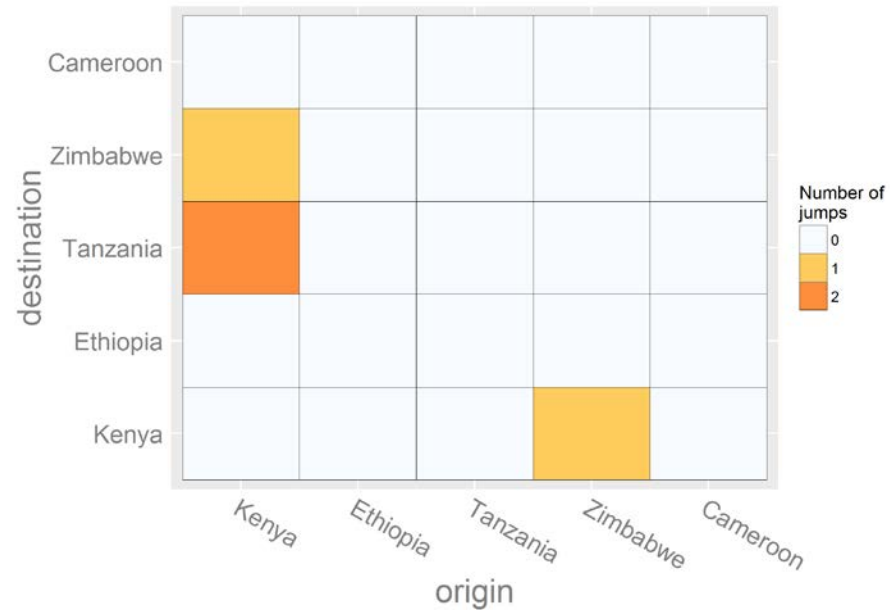
Country

- Cameroon
- Ethiopia
- Kenya
- Tanzania
- Zimbabwe

Evolutionary tree



Markov jumps





## Predictors

- Different anthropological and environmental variables might affect the transmission of FMD in Africa
- Determination of the role played by the cattle density, human density and land use in the virus transmission

## GLM

- Use of a Generalised Linear Model to parameterise the rates of transmission between two locations
- Rate as a linear function of one or more predictor variables.

Transition Rate Matrix

|   | A              | B              | C              | D              |
|---|----------------|----------------|----------------|----------------|
| A | 0              | $\lambda_{ab}$ | $\lambda_{ac}$ | $\lambda_{ad}$ |
| B | $\lambda_{ba}$ | 0              | $\lambda_{bc}$ | $\lambda_{bd}$ |
| C | $\lambda_{ca}$ | $\lambda_{cb}$ | 0              | $\lambda_{cd}$ |
| D | $\lambda_{da}$ | $\lambda_{db}$ | $\lambda_{dc}$ | 0              |

For  $i=1$  to  $n$  predictors

Predictor Rate Matrix

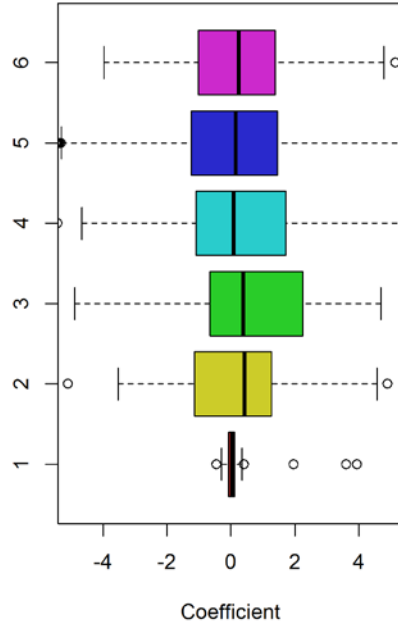
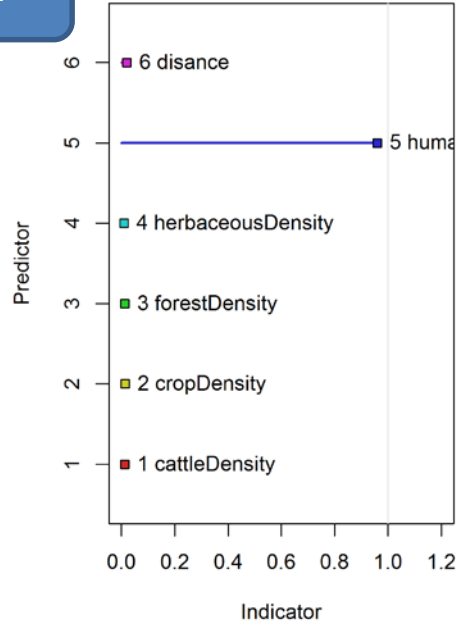
$$\log \Lambda = \sum_i \delta_i \beta_i \log R_i$$

(for each MCMC step, propose  $\beta_i$ 's and  $\delta_i$ 's)

Indicator variable, delta dirac (0 or 1)

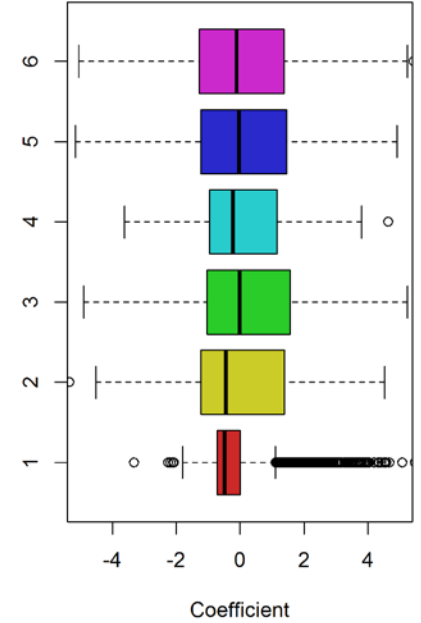
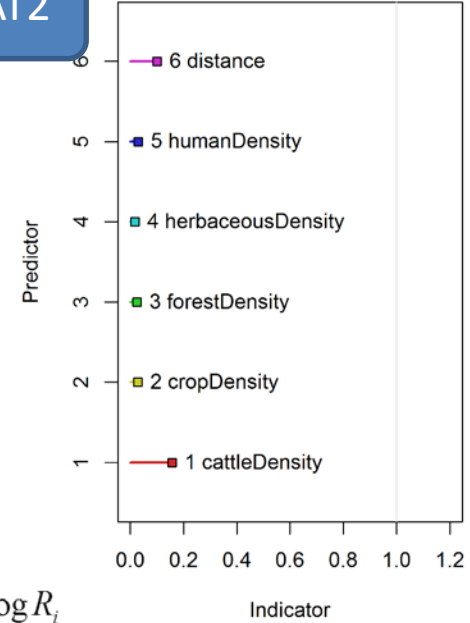
Coefficient of predictor matrix

## FMDV A



| Predictors         | BF   |
|--------------------|------|
| Cattle density     | 0,03 |
| Crop density       | 0,03 |
| Forest density     | 0,03 |
| Herbaceous density | 0,02 |
| Human density      | 58   |
| Distance           | 0,04 |

## FMDV SAT2



$$\log \Lambda = \sum_i \partial_i \beta_i \log R_i$$

↑ indicator     ↑ coefficient

| Predictors         | BF   |
|--------------------|------|
| Cattle density     | 0,4  |
| Crop density       | 0,03 |
| Forest density     | 0,06 |
| Herbaceous density | 0,06 |
| Human density      | 0,07 |
| Distance           | 0,2  |

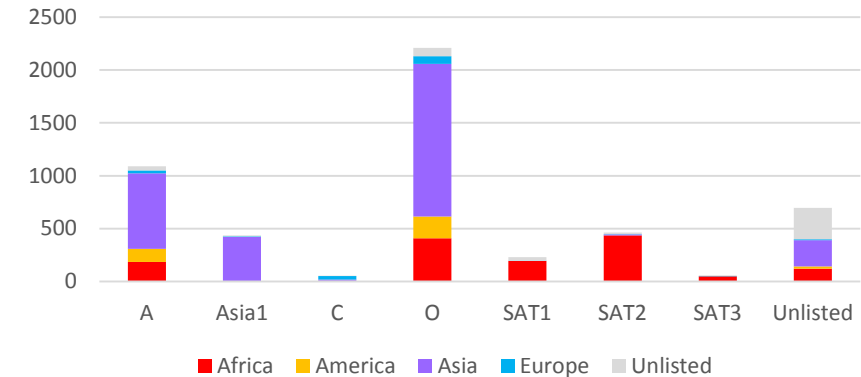
**FMDV A**

- Clear separation between the two African coasts for recent circulation
- Detection of well supported routes of transmission
- Effect of human density on the transmission

**FMDV SAT2**

- Circulates for a longer time than FMDV A
- Detection of well supported routes of transmission
- Small effect of distance and cattle density on the transmission

- Need for new FMD sequences

**Number of VP1 (1D) Sequences in Genbank**

Thanks for your attention

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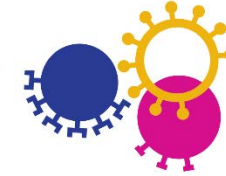
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