#### **Classification of endogenous viral elements**

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### **Endogenous viral elements (EVEs)**

Endogenous viral elements are viral nucleotidic sequences that integrated into hosts' genomes by **horizontal gene transfer**:

- **actively** through the action of a virus-encoded **integrase**: retroviruses (family *Retroviridae*) integration is an obligatory step in the virus' replication cycle
- passively by non-homologous end-joining (illegitimate recombination): *Geminiviridae* and *Caulimoviridae* sequences in plant genomes e.g – integration is not an obligatory step in the replication cycle

When integration occurs in **germline cells**, the proviral DNA is passed to future generations like a normal cellular gene and becomes **endogenous** (as opposed to transmission between individuals following conventional infection pathways)

Some endogenous viral elements are replication competent, however most of them are representative sof extinct viral taxa



## Endogenous loci as fossils representing extinct viral taxa

- Heritable, genetic loci (sequences) representing extinct viruses
- Degraded by substitutions, indels
- Subject to fixation/loss due drift & selection
- Can interleave with extant viruses in taxonomy
- Copy numbers vary :
  - up to thousands of copies in plant genomes
  - Up to hundreds of thousands of ERV copies per vertebrate genome (= billions of loci in the databases)
- 4 of 7 current *Caulimoviridae* genera represented
- 6 of 7 *Retroviridae* genera represented (ERV related to deltaretroviruses have not been discovered, yet)



### EVEs provide more resolution to phylogenies and insight into viral origins - *Caulimoviridae*: Reverse Transcriptase (RT) Phylogeny

- Entire viral genomes of extinct viruses can be reconstituted in silico from endogenous sequences:
  - 76 entire or nearly full-length genomes assembled from endogenous viral sequences, corresponding to 34 distinct species defining a new genus in the family *Caulimoviridae* tentatively named Florendovirus
  - Provide access to fossil sequences of whole virus genera that are now extinct : rice florendovirus sequences are at least 1.8 Myo



Geering et al. (2014) Nature Comms Nature Communications 5: 5269



### EVEs provide more resolution to phylogenies and insight into viral origins - *Retroviridae*: Reverse Transcriptase (RT) Phylogeny





# Why it is important to tackle the issue of the taxonomy of EVEs

- To avoid confusion that would result from unorganized naming, such as different names being given to similar sequences
- Need to provide single governing rules guiding how sequences should be named
- To prevent taxonomic terms such as family being used to describe different sequences that are no more divergent than members of a virus genus
- Need to include EVEs in the taxonomy



# Why it is important to tackle the issue of the taxonomy of EVEs



#### Lucy

Two nomenclatures:

- As a fossil specimen AL 288-1
- As evidence for an extinct hominin: *Australopithecus afarensis*



### Points for further discussion

- Endogenous loci are "fossils" representing extinct viral taxa
- Need to be incorporated into viral taxonomy just as real fossils are used to classify extinct organisms
- Incorporation of ERV and EVE into taxonomy provides more resolution to phylogenies, insight into viral origins, natural history and macroevolution
- Endogenous loci require a nomenclature that simultaneously reflects their viral origins AND their existence as genetic loci
- Additionally, these provide evidence for extinct species which will require a nomenclature and incorporation into viral taxonomy
- ERV data are vast, and functional nomenclature is urgently needed and should be in place to facilitate incorporation into taxonomic proposals

