# Emergence of Treatment-Resistant Infectious HIV after Genome-Directed Antiviral Endonuclease Therapy

<u>Daniel Stone</u>, Harshana S De Silva Feelixge, Harlan L Pietz, Pavitra Roychoudhury, Joshua T Schiffer, Martine Aubert, Keith R Jerome



# **Summary for Community**

## **Scientific Hypothesis:**

Enzymes that cut and mutate HIV DNA specifically can inactivate HIV

## **Key Findings:**

- During enzyme treatment cleavage resistant and infectious HIV emerges
- Repeated treatment with additional enzymes can negate treatment resistance

## Importance:

 Demonstrates that combination (multiple enzyme) therapies can be effective as DNA cleavage-based HIV treatments

### **Relation to Cure:**

 Disruption/mutation of the HIV genome could be used to permanently inactivate infected cells (prevent virus reactivation and provide functional cure)

### Why should we be excited?

 We can avoid the problem of resistance during the development of new DNAdirected HIV therapies