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Title: The roles of adenoviral vectors and donor DNA structures on genome editing

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The roles of adenoviral vectors and donor DNA structures on genome editing

- 1. Contrary to direct and spaced inverted repeats, nonspaced inverted repeats are intrinsically recombinogenic motifs in mammalian cells lending experimental support to their role in genome dynamics in higher eukaryotes. *This thesis*
- 2. Adenoviral, but not lentiviral, vectors constitute a valuable TALEN gene delivery platform. *This thesis*
- 3. Hitherto, the pursuit and optimization of high-fidelity donor DNA templates did not get the same emphasis as that devoted to improving the specificity of designer nuclease platforms. *This thesis*
- 4. Viral vectors constitute attractive gene delivery vehicles owing to their efficient transduction of a wide range of cell types. *This thesis*
- 5. Gene therapy cured patients with inherited immune deficiencies. *M. Cavazzana-Calvo et al. 2000. Science 288, p. 699-672; Aiuti et al. 2002. Science 296, p. 2410-2413*
- 6. The combination of high nuclease activity with reduced cytotoxicity and the simple design process marks TALENs as a key technology platform for targeted modifications of complex genomes. *C. Mussolino et al. 2011. Nucleic Acids Res. 39, p. 9283-9293*
- 7. The observed relative survival advantage of the gene-modified cells during treatment interruption suggests that genome editing at the CCR5 locus confers a selective advantage to CD4 T cells in patients infected with HIV. *Tebas P et al. 2014. N. Engl. J. Med. 370, p. 901-910*
- 8. In principle, gene therapy is simple: putting corrective genetic material into cells alleviates the symptoms of disease. In practice, considerable obstacles have emerged. *Verma and Somia. 1997. Nature. 389, p. 239-242*
- 9. Don't ever let someone tell you, you can't do something. Will Smith, The Pursuit of Happyness, 2006
- 10. Men at some time are masters of their fates: The fault, dear Brutus, is not in our stars, But in ourselves. Shakespeare, Julius Caesar Act 1, scene 2, 135–141
- 11. The difference between the impossible and the possible is merely a measure of man's determination. *Captain James Thain, pilot of the plane in the 1958 Munich air disaster*
- 12. Biology is the study of complicated things that have the appearance of having been designed with a purpose. *Richard Dawkins, The Blind Watchmaker, 1986*