

iBioIC-funded postdoc position starting August 1st 2018

This is a unique opportunity to conduct an exciting biomedical research project jointly between the University of Edinburgh and Ingenza, a biotech company based just outside the city of Edinburgh.

Many therapeutic proteins must be glycosylated to be clinically efficacious. An important example is complement factor H, a 155-kDa glycoprotein with 40 disulfide bonds that could be used to treat age-related macular degeneration (AMD). The dry form of advanced AMD is a leading cause of incurable sight loss amongst elderly populations.

It is hard to produce complement factor H and other large disulphide-rich glycoproteins in the quantities required for the pharmaceutical market place. For this purpose, yeast cells offer an attractive alternative to the traditional choice of mammalian cell culture. For example, *Pichia pastoris* grows on simple media to high densities and secretes large amounts of glycosylated recombinant proteins. However, there is a major challenge to be overcome. The pathways in *P. pastoris* responsible for glycosylation must be re-engineered so that recombinant proteins intended for human use are decorated with human-type (rather than yeast-type) glycans.

Advances in understanding of protein glycosylation and in glycan characterisation, as well as in DNA synthesis and assembly technologies, make possible a synthetic biology-based approach to re-purposing *P. pastoris* as a platform for producing humanised glycoproteins. That is the goal of this project, which is funded by the iBioIC “Accelerating Impact” programme (for 12 months in the first instance).

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