

Spatial reorganization of proopiomelanocortin (POMC)-expressing neurons in the arcuate nucleus of POMC-EGFP mice resistant or prone to obesity

Catherine Chaumontet¹, Mathilde Guillaumin², Jong Hoon Lee³, Marion Soto, Nicolas Darcel¹, Daniel Tomé¹ ¹ UMR 914, INRA-AgroParisTech, Paris, France ² Ecole Polytéchnique ³ AgroParisTech

CONTEXT

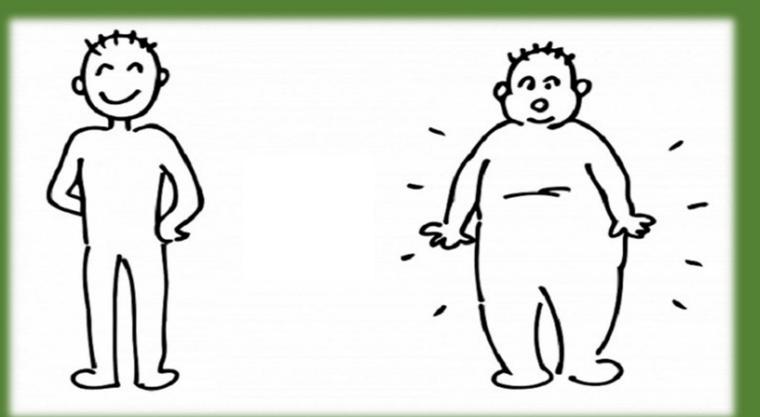
(Aged 2 weeks)

W₀

DEXA

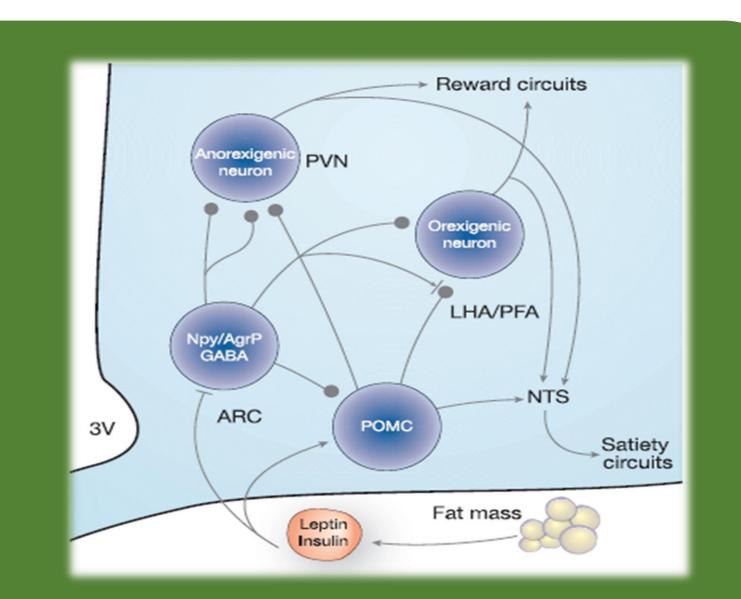
W1

Not all people exposed to a Western pattern diet become obese



(Obesity-resistant) (Obesity-prone)

Hypothalamic neural pathways involved energy homeostasis including neurons originating from the ARC that express NPY/AgRP which stimulate food intake, or POMC which are anorexigenic neuropeptides



(Morton et al 2006)

→ Do OP and OR mice have different densities and spatial distributions of POMC neurons in the ARC? Measure of spatial distribution of POMC neurons in the ARC of these two groups of mice, using 3D reconstruction and statistical mapping of POMC-neuron densities.

Experimental procedure -0.14mm to Bregma -2.90mm* coloration DAPI Control group(until aged 11 weeks) 4 OP 4 OR **Experiment 1:** 6 C57BL/6J POMC-EGFP Slicing Cryostat mice for control group Brain slicing (Aged 9 weeks) Brain slicing ARC Software: Free-D, developed by **Modeling and Digital Imaging team** gain(g) of the Institut Jean-Pierre Bourgin, INRA Versailles, France. **→** OP **Experiment 2: High Fat High Protein diet** 16 mice for OP/OR selection Habituation until aged 13 weeks Model normalisation and calibration

DEXA= Dual energy X-Ray absorptiometry

W11 7

Selection of OP/OR mice

Results Selection of OP/OR mice Body weight gain and fat mass gain of OP and OR mice → OP mice gain more weight and fat than OR mice Average daily (A) or cumulated (B) food intake of OP and OR mice → OP mice consume greater amounts of HFHP than OR mice → this leads to greater total energy intake for OP mice

W5

DEXA

W9

DEXA

POMC Neuron density in the ARC Number of POMC neurons per slice Total number of POMC neurons Control → The total number of POMC neurons are not significantly different. However, there are differences in distributions along the rostral-caudal axis Spatial distribution of POMC neurons (orange) and OR(blue) mice **Control** mice Dorsal side Caudal side Dorsal side Caudal side Rostral side Ventral side Rostral side Ventral side → 3D analyses bring to light structural differences between the three groups.

Image segmentation and repositioning

3D reconstruction of the

third ventricle and POMC neurons

CONCLUSION

Whereas global density of POMC neurons in the ARC do not seem to be influenced by an HFHP regime, results indicate differences between OP, OR and Control mice in spatial distribution and local density of these neurons. Control mice seem to have similar structures to OP mice, but further statistic comparisons must be made.

The next step is to investigate molecular mechanisms behind these spatial differences...