

XACT, a long non-coding transcript coating the active X chromosome in human pluripotent cells

Céline Vallot, Christophe Huret, Yann Leseqque, Alissa Resch, Nouf ssa Oudrhiri, Annelise Bennaceur, Laurent Duret, Claire Rougeulle

► **To cite this version:**

Céline Vallot, Christophe Huret, Yann Leseqque, Alissa Resch, Nouf ssa Oudrhiri, et al.. XACT, a long non-coding transcript coating the active X chromosome in human pluripotent cells. *Epigenetics and Chromatin: Interactions and processes*, Mar 2013, Boston, MA, United States. 6 (Suppl 1), pp.O33, 2013, <10.1186/1756-8935-6-S1-O33>. <inserm-00801773>

HAL Id: inserm-00801773

<http://www.hal.inserm.fr/inserm-00801773>

Submitted on 18 Mar 2013

HAL is a multi-disciplinary open access archive for the deposit and dissemination of scientific research documents, whether they are published or not. The documents may come from teaching and research institutions in France or abroad, or from public or private research centers.

L'archive ouverte pluridisciplinaire **HAL**, est destinée au dépôt et à la diffusion de documents scientifiques de niveau recherche, publiés ou non, émanant des établissements d'enseignement et de recherche français ou étrangers, des laboratoires publics ou privés.

ORAL PRESENTATION

Open Access

XACT, a long non-coding transcript coating the active X chromosome in human pluripotent cells

Céline Vallot^{1,2}, Christophe Huret², Yann Leseqque³, Alissa Resch⁴, Noufissa Oudrhiri⁵, Annelise Bennaceur⁵, Laurent Duret³, Claire Rougeulle^{1,2*}

From Epigenetics and Chromatin: Interactions and processes
Boston, MA, USA. 11-13 March 2013

X-chromosome inactivation (XCI), the dosage compensation process that equalizes X-linked gene expression between sexes, has mostly been studied in the mouse, where the central role for the non-coding RNA *Xist* in the initiation and spreading of the process was demonstrated. Although *Xist* is conserved in humans [1], very little is known concerning its regulation and function in this species. Several lines of evidence moreover suggest that different strategies have been adopted in the human to control XCI as compared to the mouse. In particular, in human pre-implantation development, *XIST* RNA coats the X chromosome(s) in both male and female embryos without inducing X-chromosome silencing [2]. This indicates that *XIST* expression and X-inactivation can be uncoupled during human embryogenesis and that other elements likely participate to the control of X chromosome activity in humans.

XCI is established early during embryonic development, and embryonic stem cells can be used to decipher the kinetics and the molecular actors of the process. Human female embryonic stem cells (hESC) can be found in different configurations regarding *XIST* expression: most female hESC have already undergone XCI but tend to spontaneously lose *XIST* expression [3]. In the course of an RNA-seq analysis of female hESC, we identified an extended and un-annotated transcribed region producing a long unspliced, likely non-coding nuclear RNA. RNA-FISH analysis reveals that this transcript is expressed from, and coats the active X chromosome. We called this transcript *XACT*, for X-active coating transcript. In female hESC in which *XIST* is repressed, *XACT* is expressed from and coats both Xs, and this correlates with significant reactivation of the inactive X chromosome. Expression of

XACT appears to be specific for pluripotent cells as its expression decreases during differentiation. Finally, we provide evidence that *XACT* is not conserved in the mouse.

In conclusion, we have identified *XACT* as the first long ncRNA that coats the active X chromosome in human. Given its expression profile and lack of conservation, it is tempting to speculate that *XACT* is involved in the peculiar control of XCI initiation in human.

Author details

¹Univ Paris Diderot, Sorbonne Paris Cité, Epigenetics and Cell Fate, Paris, France. ²CNRS, UMR7216 Epigenetics and Cell Fate, Paris, France.

³Laboratoire de Biométrie et Biologie Evolutive, UMR CNRS 5558, Université de Lyon, Université Lyon 1, Villeurbanne, France. ⁴Stem Cell Institute, UCHC, Farmington, CT, USA. ⁵INSERM U935, Université Paris Sud 11, AP-HP, Villejuif 94802, France.

Published: 18 March 2013

References

1. Brown CJ, Lafreniere RG, Powers VE, Sebastio G, Ballabio A, Pettigrew AL, Ledbetter DH, Levy E, Craig IW, Willard HF: **Localization of the X inactivation centre on the human X chromosome in Xq13.** *Nature* 1991, **34**:82-84.
2. Okamoto I, Patrat C, Thepot D, Peynot N, Faugue P, Daniel N, Diabangouaya P, Wolf JP, Renard JP, Duranthon V, Heard E: **Eutherian mammals use diverse strategies to initiate X-chromosome inactivation during development.** *Nature* 2011, **472**:370-374.
3. Makhlof M, Rougeulle C: **Linking X chromosome inactivation to pluripotency: Necessity or Fate?** *Trend Mol Med* 2011, **17**:326-336.

doi:10.1186/1756-8935-6-S1-O33

Cite this article as: Vallot et al: *XACT*, a long non-coding transcript coating the active X chromosome in human pluripotent cells. *Epigenetics & Chromatin* 2013 **6**(Suppl 1):O33.

¹Univ Paris Diderot, Sorbonne Paris Cité, Epigenetics and Cell Fate, Paris, France

Full list of author information is available at the end of the article