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ZSCAN 4⁺ mouse embryonic stem cells have an oxidative and flexible metabolic profile

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Abstract

Cultured mouse embryonic stem cells are a heterogeneous population with diverse differentiation potential. In particular, the subpopulation marked by Zscan4 expression has high stem cell potency and shares with 2 cell stage preimplantation embryos both genetic and epigenetic mechanisms that orchestrate zygotic genome activation. Although embryonic *de novo* genome activation is known to rely on metabolites, a more extensive metabolic characterization is missing. Here we analyze the Zscan4⁺ mouse stem cell metabolic phenotype associated with pluripotency maintenance and cell reprogramming. We show that Zscan4⁺ cells have an oxidative and adaptable metabolism, which, on one hand, fuels a high bioenergetic demand and, on the other hand, provides intermediate metabolites for epigenetic reprogramming. Our findings enhance our understanding of the metastable Zscan4⁺ stem cell state with potential applications in regenerative

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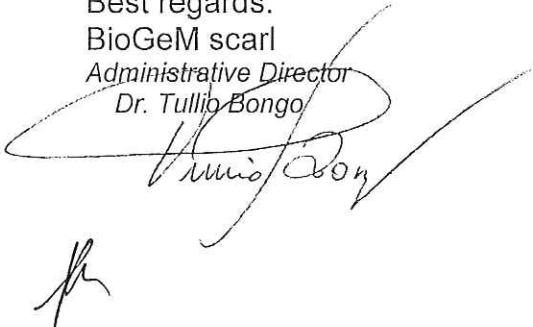
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