

ASSOCIATION OF PLASMINOGEN ACTIVATORS WITH CUMULUS EXPANSION AND OOCYTE MATURATION

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Bidirectional intercellular communications between the oocyte and their surrounding cumulus cells are essential for the oocyte developmental competence. Previous studies showed that plasminogen activators (PAs), i.e., urokinase-type (PLAU) and tissue-type (PLAT) PA, may involve in the process of oocyte maturation. How cumulus cell expressed PAs affect cumulus expansion and oocyte maturation remains to be determined. Here, we evaluate the expression of PLAT in cumulus cells of mature and immature human oocytes and use the mouse model to study the effect of the enzymatic activity of PLAU on cumulus expansion during oocyte maturation in vitro. PLAT, the human cumulus cells predominantly expressed PA, was differentially expressed between cumulus cells of immature and mature human oocytes as determined by the quantitative real-time polymerase chain reaction. To assess the effect of the enzymatic activity of PLAU on oocyte in vitro maturation (IVM), two PLAU-specific inhibitors, i.e., 4-chlorophenyl guanidine hydrochloride (4CGH) and UK122, were added before oocyte IVM. PLAU specific inhibitors were dose-dependently inhibited mouse cumulus-oocyte complex expansion and the subsequent oocyte maturation. The dynamics of cumulus expansion recorded by a time-lapse image analyzer revealed that the inhibition of cumulus expansion by 4CGH was first detected at 6 h during IVM. Increasing fetal bovine serum concentration in culture medium from 10% to 20% reversed the suppression of 4CGH in a dose-dependent manner. Replacement of medium serum to the fetuin protein showed a similar inhibitory effect of the 4CGH on cumulus expansion and oocyte maturation. Taken together, the PA may involve in the process of oocyte maturation. The enzymatic activity of PLAU in mouse oocyte in vitro maturation is primarily contributed by the endogenous expression of cumulus cells though it also derived from serum supplementation to the IVM medium.