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Involvement of A₁ Adenosine Receptors in the Acquisition of Fertilizing Capacity

ALBA MINELLI^{*}, LAVINIA LIGUORI^{*}, ILARIA BELLAZZA^{*}, ROBERTA MANNUCCI[†], BJÖRN JOHANSSON[‡] AND BERTIL B. FREDHOLM[‡]

From the ^{*} Dipartimento di Scienze Biochimiche e Biotecnologie Molecolari, Sezione Biochimica Cellulare, Perugia, Italy; [†] Dipartimento di Medicina Interna e Scienze Oncologiche, Policlinico Monteluce Perugia, Italy; and [‡] Department of Physiology and Pharmacology, Section of Molecular Neuropharmacology, Karolinska Institutet, Stockholm, Sweden.

Correspondence to: Dr Alba Minelli, Dipartimento Scienze Biochimiche e Biotecnologie Molecolari, Sezione Biochimica Cellulare, Via del Giochetto, 06123 Perugia, Italy.

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a two-step process: capacitation followed by acrosome reaction. The biochemical and biophysical modifications occurring in vivo in the female reproductive tract can be reproduced in vitro, and previous studies have suggested a capacitative role for adenosine A_1 receptor (A_1R). Mice with a targeted disruption of the Adora 1 gene (A_1R –/- mice) provide a useful model for better understanding the role of the A_1R in fertility. Murine spermatozoa express A_1R in the head, neck, midpiece region, and tail. The number of capacitated spermatozoa incubated in human tubal fluid was significantly reduced in A_1R –/- compared with A_1R +/+ and A_1R +/- spermatozoa. The difference between $A_1 R$ +/+ and A_1R -/- mouse spermatozoa was mainly in the time necessary to reach the maximum percentage of capacitation. A_1R +/+ murine sperm obtained the full state of capacitation within 90 minutes whereas A_1R –/- sperm required 240 minutes. Caffeine, a known antagonist of A_1 and A_{2A} adenosine receptors, lowered the number of capacitated sperm and affected the time of capacitation in a dose-dependent manner, mimicking the effects of the lack of A_1 receptors. Although number, motility, and viability of A_1R –/- murine sperm was not significantly different from A_1R +/+ mouse spermatozoa, a significant reduction of the number of pups produced by A_1R -/- male mice suggests that A_1 receptors must be fully operative to accomplish the optimal degree of capacitation and thereby fertilization.

Key words: A₁ adenosine receptors KO mice, A₁ adenosine receptors mouse sperm localization, capacitation, caffeine, fertility

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