C-C Motif Chemokine Ligand 2 Regulates Prostaglandin Synthesis and Embryo Attachment of the Bovine Endometrium during Implantation

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Embryo Attachment of the Bovine Endometrium during Implantation

(着床期ウシ子宮内膜における CC モチーフケモカインリガンド 2 によるプロスタグランジン合

成と胚接着の制御)

Category: Koh

Thesis Summary

C-C motif chemokine ligand 2 (CCL2), which regulates immune cells, has been

reported to be expressed in the bovine endometrium during pregnancy. However, the

details of how CCL2 is involved in the implantation mechanism of bovine embryos are

still unclear. The purpose of this study is to analyze the expression pattern and

functional properties of CCL2 in the bovine endometrium and embryos.

The qPCR analysis of the tissue showed that not only the amount of CCL2, but also

CCL8 and CXCL10 were high at the implantation stage. The amount of CCL2 was

significantly high in IFNa treated bovine endometrial stromal (BES) cells in vitro. In

bovine endometrial epithelial (BEE) cells, however, the amount of CCL8 and CXCL10

were significantly high in the treatment group, but not for CCL2. The mRNA of each

chemokine receptors (CCR1, CCR2 and CXCR3) was detected in the endometrial

tissues and cells by RT-PCR. Cellular proliferation of BEE and BES significantly

increased by the CCL2 treatment. The amount of prostaglandin (PG) E2 synthases,

PGES1 and PGES2, and PGF2 alpha synthase, AKR1C4, were high at the implantation

stage compared with luteal stage. The amount of PGES2 and AKR1B1 were

significantly increased by CCL2 treatment does-dependently in BEE. In BES, on the

other hand, the amount of PGES3, AKR1A1 and AKR1C4 were increased by CCL2

treatment. The qPCR analysis of the tissue showed that there were no differences in the

amount of PGs transporter transcripts (MRP4 and PGT) between the luteal and

implantation stages. The amount of *MRP4* and *PGT* were significantly high in CCL2 treated bovine endometrial epithelial (BEE) and stromal (BES) cells *in vitro*. The mRNA of chemokine receptors (*CCR1*, *CCR2* and *CXCR3*) were detected in the bovine trophoblastic cells derived from the blastocyst (BT) by RT-PCR. The amount of *PCNA* and *IFNt* were significantly high in the BT treated with CCL2 compared to the control. CCL2 significantly increased the attachment rate of BT vesicles to BEE in *in vitro* co-culture system. The amount of *OPN* increased in BEE, and *ICAM-1* increased both in BEE and BT by CCL2 treatment.

These results indicate that CCL2, which expression increased in bovine endometrium during implantation by embryonic factors, has the potential to regulate the synthesis and circulation of PGs in the endometrium and the embryo growth. In addition, CCL2 has a possibility of regulating the process of bovine embryo attachment to the endometrium by modulation of binding molecules.