

部門公開セミナー

共催:新学術領域研究「配偶子幹細胞制御機構」

Spermatogenesis – novel proteins and structures

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For over a decade, our laboratory has been using knockout and knockin transgenic mouse studies to define TGF-beta superfamily signaling pathways, study ovarian and testicular cancers, and understand the functions of germ cell-expressed genes. To understand reproduction and cancer *in vivo*, we have produced over 50 transgenic knockout models including mice lacking TGF-beta superfamily ligands [e.g., growth differentiation factor 9 (GDF9), bone morphogenetic protein 15 (BMP15), activins, and inhibins]. To isolate novel gonad-expressed genes, we have used multiple strategies including proteomics, degenerate PCR, cDNA sequencing, *in silico* (electronic database) subtraction, PCR-based subtraction, genome database mining, and GeneChip technology. In the process of these studies, we have identified over 20 novel and gonad-expressed genes. Because fertility cannot be evaluated in a test tube, we have generated knockout models to define the *in vivo* functions of many of these novel genes. Our studies have identified genes that play key roles in the ovary (e.g., GDF9, BMP15, and pentraxin), in the testis (e.g., Tex14 and Klhl10), as maternal effect genes (e.g., zygote arrest 1 and nucleoplasmin 2), or as tumor suppressors (e.g., inhibins). These functional genomics approaches are helping to define mechanisms of reproductive function and will continue to aid in the identification of candidate genes that are potential contraceptive targets and/or mutated in men and women with infertility or cancer.