

fluorescent dye that stained only the cortical cells at the beginning of the culture were located in the testis cord of the left testis.

Conclusions: The results suggested the presence of Sertoli cells derived from the cortical cells expressing DMRT1 in the left testis after the onset of testicular development.

### P3-52.

#### Functions of estrogen receptor $\alpha$ in different sub-cellular locations

(専攻生: 人体構造学)

○内田 俊輔

(人体構造学)

李 忠連、永堀 健太、河田 晋一  
表原 拓也、宮宗 秀伸、伊藤 正裕

<Introduction> Estrogen receptor alpha (ER $\alpha$ ) plays crucial roles in control of proliferation, differentiation, migration, and physical functions of uterine endometrial cells. It is observed that ER $\alpha$  is located not only in cell nucleus, but also in cytoplasm and membrane, and such subcellular locations varies under certain circumstances. A lot of studies indicate that PI3K-AKT is one of the major signal transduction pathways down-stream of ER $\alpha$ . <Methods> To further discern functions of ER $\alpha$ , it was allocated to different subcellular locations in ER $\alpha$ -negative Ishikawa cell, using genetic recombination and permanent transfection techniques. Cells with ER $\alpha$  expressed only at cell membrane, cytoplasm, or nucleus were cloned, and these with similar amount of expression in ER $\alpha$  were further selected for experiments. These cells were observed and analyzed with Annexing V staining, scratch assays, immuno-histological staining, FACS, and other related techniques.

<Discussion> The results show that ER $\alpha$  of different subcellular locations, together with some of its down-stream signaling pathways, regulates cell size and migration with different intensity. It was observed that cells with membranous ER $\alpha$  were significantly faster in migration than control, and such cells were significantly bigger in size than the others. Further analysis showed that PI3K-AKT-mTOR pathway down-stream of ER $\alpha$  is

closely involved in regulation of cell size. The present study indicates that ER $\alpha$  in different subcellular locations possesses respective effects on some crucial cellular events.

### P3-53.

#### AAV-mediated miRNA-29b delivery suppress renal fibrosis

(社会人大学院博士課程 2年腎臓内科学)

○齋藤 優

(腎臓内科)

菅野 義彦

(分子病理学)

大野慎一郎、原田裕一郎、老川 桂生

黒田 雅彦

Renal fibrosis, characterized as the accumulation of excess extracellular matrix (ECM) on the kidney parenchyma, is the principal process underlying the progression of chronic kidney disease (CKD) to end-stage renal disease (ESRD), independently of the primary renal disease which causes the original kidney injury. MicroRNAs (miRNAs) are endogenous short non-coding RNAs that regulate post-transcriptional gene expression. Recent studies have shown that miRNA-29b protects kidney from renal fibrosis by suppressing the deposition of ECM and preventing epithelial-to-mesenchymal transition (EMT). However, an efficient method of gene transfer into the kidney has not been established. Here, we report a kidney-targeted gene delivery using recombinant adeno-associated virus (rAAV) vectors delivered by injection into the renal pelvis in mice. We also discuss that which serotypes of AAV vector is suitable for kidney-targeted gene delivery and the optimized second structure of miRNA-29b to be cleaved into the mature form.