

# Fetal Allograft Survival in Immunocompetent Recipients Is Age Dependent and Organ Specific

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This study explores whether fetal allograft survival is age dependent and organ specific. Fetal rat tissue (renal, gonadal, hepatic) from the third trimester of gestation (days 15–21) was transplanted into 306 outbred adult rats for 10–30 days. Grafts were studied by morphometric and histologic analysis. Ten days after implantation, renal tissue ( $N = 75$ ) from late gestation (days 19–21) showed no increase in size. In contrast, 17-day fetal grafts ( $N = 20$ ) grew  $6.8 \pm 3.4$  times,\* while 15-day fetal grafts ( $N = 28$ ) grew  $17.5 \pm 6.1$ \* times. (The symbol "\*" indicates  $p < 0.05$ , compared to original size.) Twenty days after implantation, these 15-day fetal grafts ( $N = 20$ ) grew  $48.8 \pm 17.7$ \* times. Ten days after grafting, the younger fetal tissue showed excellent maturation of renal elements and no sign of rejection; older fetal grafts had poor renal architecture and a dense lymphocytic infiltrate. The 15-day fetal gonadal tissue ( $N = 18$ ) showed a moderate  $10.6 \pm 3.2$ \* increase in size while the 15-day hepatic grafts ( $N = 16$ ) were regularly rejected within 10 days. Selected fetal allografts from early in the third trimester can not only survive but can grow and mature in an immunocompetent recipient. This fetal graft growth appears to be both age dependent and organ specific. The use of fetal organs may broaden the potential pool for transplantation. However, further studies are needed to define the ontogeny of graft acceptance.

**I**N AN ATTEMPT to broaden the pool of available organs for transplantation and to understand better the mechanisms of graft rejection, this laboratory has embarked on a program to investigate the antigenicity of fetal donor tissue. It is well known that the immunocompetent adult promptly rejects allogeneic adult tissue soon after transplantation.<sup>1</sup> Studies from other laboratories using fetal small intestinal or pancreatic tissue from late in

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gestation showed prompt rejection in allogeneic adult recipients.<sup>2,3</sup> In contrast, we have demonstrated that fetal kidney and adrenal tissue from early in the third trimester survived and grew when transplanted into adult allogeneic recipients.<sup>4</sup> These seemingly divergent results prompted us to ask whether fetal graft growth might be both age dependent and organ specific. To this end, we varied the ontogeny and the organ of the fetal grafts that were implanted into adult allogeneic animals.

## Materials and Methods

### *Animals*

Outbred Sprague-Dawley rats were obtained from Holtzman Laboratories, Madison, Wisconsin. All recipient animals were adult females over 10 weeks of age and weighed 225–250 g. Timed pregnant mothers were shipped between 11 and 13 days of pregnancy. A total of 306 animals received grafts.

### *Graft Tissue*

The graft tissue consisted of renal tissue harvested from adults (Group I) or fetal tissue: renal (Group II), gonadal (Group III), or hepatic (Group IV). Adult donor rats underwent cervical dislocation, and their kidneys were rapidly excised, placed in Dulbecco's Modified Eagle's Medium at 4°C, and divided under a dissecting microscope into fragments less than 1 mm in diameter. Fetuses were delivered by cesarean section from timed pregnant mothers on the fifteenth to twenty-first day of gestation. The liver, kidneys, and gonadal tissue from successive days in gestation were excised, placed in 4°C culture medium, and

Presented at the 106th Annual Meeting of the American Surgical Association, Hot Springs, Virginia, April 24–26, 1986.

This investigation was supported by PHS Grant #K08-CA01023-01, awarded by the National Cancer Institute, Clinical Investigator Award (RPF); Directed Giving Award-Johnson & Johnson (PKD); and the Eleanor B. Pillsbury Fellowship—University of Illinois Hospital (MBS).

Reprint requests: Pediatric Surgical Research Laboratory, Massachusetts General Hospital, Boston, MA 02114.

Submitted for publication: April 28, 1986.

# Mechanisms Of Fetal Allograft Survival

**J. Larry Jameson, Leslie J. De Groot**



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