

Article

Integrating Transcriptomic and ChIP-Seq Reveals Important Regulatory Regions Modulating Gene Expression in Myometrium during Implantation in Pigs

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Abstract: The myometrium is the outer layer of the uterus. Its contraction and steroidogenic activities are required for embryo implantation. However, the molecular mechanisms underlying its functions remain unknown in pigs. The myometrium includes the inner circular muscle (CM) and the outer longitudinal muscle (LM) layers. In this study, we collected the CM and LM samples from the mesometrial side (named M) of the uterus on days 12 (pre-implantation stage) and 15 (implantation stage) of pregnancy and day 15 of the estrous cycle. The transcriptomic results revealed distinct differences between the uterine CM and LM layers in early pregnancy: the genes expressed in the LM layer were mainly related to contraction pathways, whereas the transcriptional signatures in the CM layer on day 15 of pregnancy were primarily involved in the immune response processes. Subsequent comparisons in the CM layer between pregnant and cyclic gilts show that the transcriptional signatures of the CM layer are implantation-dependent. Next, we investigated the genome-wide profiling of histone H3 lysine 27 acetylation (H3K27ac) and histone H3 lysine 4 trimethylation (H3K4me3) in pig uterine CM and LM layers. The genomic regions that had transcriptional activity and were associated with the expression of genes in the two layers were characterized. Taken together, the regulatory regions identified in the study may contribute to modulating the gene expression in pig uterine CM and LM layers during implantation.

Keywords: pigs; implantation; myometrium; regulatory regions; H3K27ac; H3K4me3

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1. Introduction

Litter size is one of the most economic traits in swine production. Multiple factors and processes determine the pregnancy outcome. Success in implantation and embryo development is required to achieve a large litter size. The uterus plays a fundamental role in implantation and embryo development during pregnancy. Structurally, the uterus consists of three layers: the endometrium, myometrium, and perimetrium [1]. An endometrium is where the embryo implants and resides during pregnancy. Thus, its role in embryo implantation has been studied intensively and well-determined [2–6]. The myometrium consists of two layers of smooth muscle. Its primary function is to generate uterine contraction and steroidogenic activity, which are also required for implantation [7–9].

Much progress has been made toward understanding the physiological mechanisms of myometrial activity in implantation. In polytocous species such as rabbits, mice, rats, and pigs, it is ideal that embryos can implant evenly along the uterine horns. Thus, intrauterine embryo spacing is a common event during pre-implantation. The mechanical force generated by the myometrial activity of the uterus is essential in the migration of embryos in the uterus [9–11]. In addition, the myometrium also plays an endocrine role

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Progesterone Its Regulatory Effect On The Myometrium Ciba Foundation Study:

Progesterone: Its Regulatory Effect on the Myometrium Brenda Muriel Schofield, 1969 **Progesterone** Ciba Foundation, 1969 **Physiology and Genetics of Reproduction** Elsimar Coutinho, 2014-11-14 **Current Catalog** National Library of Medicine (U.S.), 1979 First multi year cumulation covers six years 1965 70 *Medical Journal of Australia*, 1914 *The Fetus and Birth* Julie Knight, Maeve O'Connor, 2009-09-16 The Novartis Foundation Series is a popular collection of the proceedings from Novartis Foundation Symposia in which groups of leading scientists from a range of topics across biology chemistry and medicine assembled to present papers and discuss results The Novartis Foundation originally known as the Ciba Foundation is well known to scientists and clinicians around the world **Perinatal Medicine** P. J. Huntingford, R. W. Beard, F. E. Hytten, J. W. Scopes, 1971-09-14 **Pregnancy and Parturition** T. Zakar, 1996-06-25 One of the mysteries of mammalian reproduction is the physiologic process that determines the length of gestation The proper timing of birth ensures that the young individual is sufficiently developed to survive and adapt in the extrauterine environment and that the mother is capable to provide nutrition and protection to the newborn This volume summarizes new knowledge obtained by many researchers seeking to unravel the compile mechanisms that contribute to the maintenance and termination of pregnancy The most important common goal of these efforts is to reduce the incidence of preterm birth which is the leading cause of perinatal morbidity and mortality in numerous countries Separate chapters are devoted to the best studied animal models of parturition In sheep the fetus is in control of the timing of its own birth while in avian species oviposition is evidently determined by the female laying the feritlized egg In humans and non human primates the roles of the fetus and the mother are more balanced and involve a complicated and poorly understood interplay between the mother the fetus and the placenta Some major aspects of these interactions such as trophoblast function myometrial contractility and the endocrine paracrine systems are discussed in further chapters **Progesterone**, 1969 **Preventing Low Birthweight** Institute of Medicine, Division of Health Promotion and Disease Prevention, Committee to Study the Prevention of Low Birthweight, 1985-01-01 Despite recent declines in infant mortality the rates of low birthweight deliveries in the United States continue to be high Part I of this volume defines the significance of the problems presents current data on risk factors and etiology and reviews recent state and national trends in the incidence of low birthweight among various groups Part II describes the preventive approaches found most desirable and considers their costs Research needs are discussed throughout the volume **National Library of Medicine Current Catalog** National Library of Medicine (U.S.), 1965 **Interactions Between Steroids and Spasmogens on the Myometrium of the Ewe** Michael John Carrick, 1973 **National Union Catalog**, 1973 Includes entries for maps and atlases **Biology of the Uterus** Ralph Wynn, 2013-06-29 In the decade following the publication of the first edition of *Cellular Biology of the Uterus* advances in this field have been so rapid as to require not merely a revision of the earlier text but an essentially new volume Even the title of

the book has been changed to Biology of the Uterus to reflect the incorporation of more material based on classical anatomy and physiology This histological and embryological information provides a necessary though often lacking background for the protein chemist and molecular biologist and a bridge between biochemistry and biophysics on the one hand and clinical medicine on the other Thus major practical problems in human reproduction such as the mode of action of contraceptive agents and the cause of the initiation of labor may be approached on a firm scientific footing This text deals primarily with the biology of the uterus itself comparative and human rather than with placentation or pregnancy and as such is a synthesis of data derived from many techniques conventional and modern Inasmuch as it is clearly beyond the competence of anyone scientist to prepare such a text on the basis of personal knowledge and experience the aid of distinguished biologists from this country and abroad was enlisted All of these authors acknowledged experts in their respective fields agreed to extensive revision of their chapters or preparation of entirely new contributions

Obstetric Therapeutics Denis Frank
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