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## Human blastocyst formation in vitro

RNA-Protein Binding IGF2BP2/IMP2 is a critical maternal activator in the early zygotic genome of activation. Liu HB, Muhammad T, Guo Y, Li MJ, Sha QQ, Zhang CX, Liu H, Zhao SG, Zhao H, Zhang H, Du YZ, Sun K, Liu K, Lu G, Guo XJ, Sha J, Fan HY, Gao F, Chen ZJ, Liu HB, et al. *Adv Sci (Weinhl)*. 2019 May 24;6(15):1900295. doi: 10.1002/adv.201900295. eCollection 2019 Aug 7. *Adv Sci (Weinhl)*. 2019. PMID: 31406667 Free PMC article. Human Blastocyst (day 5)[1] (Greek, blastos = kal + cistos = cavity) or blastula, a term used to describe hollow cell mass formed in early development. The blastocist consists of cells that form a layer of trophoctoderma (TE, trophoblast), internal cell mass (ICM, embryonic explosion) and blastogel (fluid-filled cavity). The internal cell mass will form the entire embryo and is the source of the true stem stem cells, which can form all types of cells within the embryo. In mammals, trofektoderm will form key cells (trophoblasts) of the fetal component of the placenta. In humans, the development phase of the blastocyst occurs in the first and second weeks after fertilization (week 3 and 4 GA) and is initially described as the Carnegie Phase 3. This phase is followed by blastocyst ejection and implantation. One of the first key development decisions in the moruli for the development of blastocyst is the fate of TE cells or ICM. Blastomers genome divergenca[4] Mouse development blastocysts[5] Some recent findings Computing data analysis of single-cell transcriptic elucidating stabilization of the Oct4 expression in the composition E3.25 mouse preimplantation [6] Our computational analysis is focused on transition 32- to 64-cell mice embryos. Embryonic day (3.25), chiju is the study u literature concentrated umozeno on nuddu for the first race second cell-fate decision. Specification of internal cell mass (ICM) to primitive endoderma (PE) i epiblasta (EPI). We analyzed single-cell (sc) data on microarray transcriptomics from E3.25 using hierarchical Optimal k-Means (HOKM) clusters, I identified two groups of ICM-cell: a group cell from an embryo with less than 34 cells (E3.25-LNCs), and another group cell from an embryo with more than 33 cells (E3.25-HNC) corresponding to two developmental phases. Although we found massive heterogeneity in ICM cells at E3.25-HNC with more than 3,800 genes with transkriptomu bifurcation, many of which are PE and EPI markers, we showed that E3.25-HNCs are not EITHER PE or EPI. Importantly, in analyzing the genes expressed differently between E3.25-LNC and E3.25-HNC, we discovered a non-autonomous mechanism based on the minimum number of four internal cell contacts in the ICM that activates Oct4 in pre-implantation brood. Oct4 is highly expressed but unstable at E3.25-LNC and stabilises at a high level at E3.25-HNC, with a high-expressed Bsg and to establish an early naial pluripotent state. Our results show that the pluripotent state that we found in ICM at E3.25-HNC is the in vivo counterpart of a new, very early pluripotent state. We compared the transcriptic profile of this in vivo E3.25-HNC pluripotent state, Together with the profile E3.25-LNC, E3.5 EPI and E4.5 EPI cell, with the profiles of all embryonic stem cells (ESC) available u GEO database from the same platform (over 600 microarrays). Physiological profile of non-differentiated bovine blastocysts derived from trophoblasts[7] Trophoctoderm blastocysts mediate early events in communication between the foetus and mothers, allowing implantation and establishment of a functional placenta. Inadequate or impaired developmental events associated with trophoblasts have a direct impact on the early survival of the foetus and successful implantation at a pivotal period corresponding to a high incidence of pregnancy losses in dairy cows. The molecular basis of bovine trophoctoderma development and signaling for the start of implantation is still not well understood. In this study, we developed methods for growing non-differentiated bovine blastocyst trophoblasts and used both transcriptomy and proteomics in early colonies to categorize and elucidate their functional characteristics. Based on absolute abundance, 9270 transcripts and 1,418 proteins were identified and analysed. We profiled a comprehensive list of growth factors, cytokines and other important factors that can effectively affect paracrine communication in uterine microsil. Functional categorisation and analysis revealed new information on structural organisation, composition of extracellular matrix, cell composition and components for adhesion, transmission networks and metabolic preferences. Our data show the fundamental physiology of bovine trofektoderma and indicate signs of a self-renewable non-differentiation state similar to the trophoblast stem cells described in other species. Four simple rules sufficient to generate mammalian blastocysts[5] We implemented experimentally reported mechanisms for polarity, cell signaling, adhesion and apoptosis as a set of developmental rules in silicone model of physically interacting cells. It is found that this model reproduces specific mutant phenotypes quantitatively and provides an explanation for the occurrence of heterogeneity without the need for an initial transcription variation. It also suggests that the fixed time point for cell competence for fibroblast growth factor (FGF)/extracellular signal regulated kinase (ERA) should determine the embryonic clock allowing certain occurrences of modification, a concept that is quantified by in vitro embryo manipulation. Based on these observations, we conclude that the minimum set of rules allows the embryo to experiment with the stochastic gene and could provide the strength needed to diversification of the regulatory network for gene preimplantation. More recent articles This table enables automated computer search of an external PubMed database using the specified text link Search Term. This search now requires a manual connection because the original PubMed extension was disabled. The list of references displayed does not reflect any editorial selection of material based on content or relevance. References also appear in this list based on the date of actual page views. References listed on the remaining content pages and related discussion pages (listed under the sub-attachments for the year of publication) include some editorial selection based on relevance and availability. More? References | Discussion page | Log Searches | 2019 References | 2020 Reference Search Term | Blastocist | Blastocoeel | Internal cellular mass | | Trophoctoderm Older Terms: These papers originally appeared in the table Some recent findings, but as this list has grown to length, they have now been shuffled down to this broken down table. See also the discussion page for other references listed by year and references to that current page. Asymmetric division of shrinking cell positioning of cell domains and fate specification[8] During the development prior to mammalian implantation, the embryo is only organized into a blastocyst consisting of an epithelial layer that encapsulates the inner mass cell (ICM), which causes all embryonic tissues. In mice, targeted cell division, acolytic polarity and actomyosine contraction contributed to the formation of ICM. However, it remains unclear how these processes work together. Here we show that asymmetric segregation of apical domains creates blastomers with different contractility, triggering their classification into internal and internal positions. Oct4 is required for lineage in the development of the internal cell mass of the blastocyst mouse[9] The Oct4 transcription factor is required in vitro for the establishment and maintenance of embryonic stem cells and for the reprogramming of somatic cells to pluripotency. In vivo, it prevents ectopic differentiation of the early embryo into trophoblast. Here we further explore the role of Oct4 in the formation of blastocyst and the specification of epiblast compared to primitive endoderm lines using conditional genetic deletion. Experiments involving mouse embryos, which are deficient for both the uterine and zygotic Oct4, suggest that this can be used to create zygote, early cleavage and activate nanog expression. Nanoga proteins are significantly elevated in the purported internal cellular mass of Oct4 null embryos, indicating an unexpected role for Oct4 in reducing Nanog levels that may be relevant for differentiation in priming differentiation during epiblastation. In-order erasure of Oct4 during the transition of the morula to blastocist interferes with the ability of internal cellular mass cells to adopt a linear-specific identity and gain a molecular profile epiblast or primitive endoderma. Sox17, a marker of primitive endoderma, is not detected by the long-term culture of such embryos, but can be solved by providing exogenous FGF4. Interestingly, functional primitive endoderm can be solved in Oct4-deficiency embryos in complementary embryos, but only if the hosts of the embryos are at the pre-blastocist stage. We conclude that decisions on the fate of cells within the internal cell mass depend on Oct4 and that Oct4 is not independent to distinguish primitive endoderma derivatives until an appropriate developmental environment is established. Non-invasive imaging of a human embryo prior to embryo activation predicts development in the blastocyst phase[10] Studies of pre-implantation of human embryo development have been reported, which correlates image analysis at times and profiling of genetic expression. By studying a large set of in vitro fertilisation (IVF) studies, success in progression to the blastocyst phase can be predicted with 84t:93% sensitivity and specificity by measuring three dynamic, non-invasive imaging parameters after day 2 after fertilisation, before the activation of the embryo genome (EGA). The expression of the blastocyst gene is associated with the potential of implantation[11] Compared to the blastocysts that caused healthy foetal development, blastocysts that do not react (negative) showed that the b3gn15 i Eomes gene was smeared, a blastocyst that resulted in spontaneous loss of pregnancy (absorption) showed a decrease in the Wnt3a i Eomes gene. FGF signal-dependent segregation of primitive endoderma and epiblastic in the mouse blastocyst[12] Primitive endoderm (PE) and epiblast (EPI) are two lines derived from the internal cell mass (ICM) of the E3.5 blastocyst. Recent studies have shown that epi and pe progenitors, which express lineano-specific transcriprial factors Nanog and Gata6 respectively, gradually occur with the development of ICM. ... In conclusion, we propose a model in which the stochastic and progressive specification of EPI and PE lineage occurs during the maturation of the blastocyst on a signal-dependent FGF/MAP. Movies Human Blastocyst Model Development Morula Model Page | Play blastocist model page | Play Movies Recent #pmid28681376# PMID28681376) of the initial composition of the morula to blastocyst, based on animal models, provides important mechanical steps: Compaction - morula blastomers packing tightly (microfilament cytoskeleton) Cleavage level - spindle direction of dividing cells (mitozta) Polarization - blastomers apico-basal (Hippo pathway, Yes-associated protein (Yap) Cavitation - blastokoeel formation with enlargement cycles and cavity collapse. epiblast cells - contact polar trophoctoderm primitive endoderm - facing the cavity Formation of human blastocyst Below table shows changes in development of human blastocyst in vitro within week 1, [13] With the designation blastist blastocist Blastocist Blastocist Blastocist Blastocist Blastocist from the pellucida (human) Model Human Blastocyst Development The next figure is from a recent study[10] using video and genetic analysis of human development in vitro within 1 week of fertilization. EGA - embryonic activation of the EUSEF genome - embryo phase specific sample, four unique embryonic phase-specific samples (1-4) Links: Image with legend Mouse Blastocyst Gen Expression General patterns of genetic expression are indicated from genomic profiling [15] red - loss of maternal mRNAs green - activation of the embryonic genome (EGA) purple - maternal gene activation (MGA) orange - continuous expression Internal cell mass Human Blastocyst (day 5)[1] The mass of the nud cells forms the inner layer of the larger cell you are with your embryoblast is a cluster of cells located on one wall of the trophoblast layer. In week 2, this mass will vary into two different layers of epiblast and hypoblast. Hypoblast (or primitive endoderma) is a transient epithelial layer facing blastocoeel, replacing it within three weeks with endoderm cell gastrulation. The epiblastic layer will form the entire embryo and within three weeks undergo gastrulation to form 3 sprouts of the layer. It also forms an epithelial layer that padding the amniotic cavity. Trophoctoderm Trophoctoderm (TE) is also called the trophoblast epithelial. It is also called the trophoblast epithelial, which is a key component of the placenta. A key early function is to transport sodium (Na+) and chloride (Cl-) ions through this layer into blastocoi. Later in week 2, this layer will differentiate into two different trophoblast layers, the cells of syncytiotrophoblast and cytotrophoblast and are key to implantation and early placenta. The early layer differentiation has been shown to be governed by the transcription factors Tead4[16] and then homeobox 2 (Cdx2) associated with Caudal. Links: trophoblast | OMIM - Tead4 | OMIM - Cdx2 Blastocoeel Mouse - blastocoeel formation[15] trophoctoderm transports Na+ and Cl-ions through this layer in blastocoeel creates an osmotic gradient of the driving fluid across this epithelial separate apical and basolateral membrane Domain Specific for transport facilitates transepitelin Na+ i transport fluid for blastocoeel formation transport commutes Na, K-adenosine triphosphatase (ATPase) u basaltic membranes trophoctoderma [17] Mishide blastocist GLUT8 term. [18] During the blastocyst phase, mammalian metabolism switches to anaerobic metabolism of glycolysis to meet the metabolic requirements of the growing blastocyst and the formation of blastocoeel. This is expected to be driven by an integral membrane protein family of glucose-relieved transporters (GLUT or SLC2A), aerobic - oxidation of lactate and pyruvate in the citric acid cycle (Krebs cycle) and oxidative phosphorylucose-converting glucose Piruvat GLUT - GLUTose Transporter (divided into 3 classes I-III) SLC2 - Solute Carrier Family 2 Glut transporter glut1 - from zygote to blastocyst. (all mammalian tissues, basal glucose withdrawal) GLUT2 and GLUT3 - from the late eight cell levels to the blastocyst. (GLUT2, beta cells of the liver and pancreas; glut3, all mammalian tissues, basal glucose uptake) GLUT4 - not expressed. (muscle and adipose tissue) GLUT8 - up-regulated in the blastocyst phase. (central nervous system and heart) (Data mainly from mouse development, expression of adult tissues shown in parentheses) Mouse study,[18] showed that GLUT8 is up-regulated after stimulation of insulin, although the recent glut8 knockout mouse shows normal early embryo development in the absence of this transporter. [19] Links: Biochemistry - glucose transporters | GLUT1 | GLUT2 | GLUT8 Blastula Cell Communication Two types of cell junctions have been identified in different regions in the development of blastocyst. The close intersections, located near the outer surface, create a leak, insulating the inside of the embryo from the external medium. The gaps allow for electrical compression of the epithelial cells that surround the fluid-filled cavity. Adhesion EM Images: GIT EPITHELIIUM EMI | GIT epithelium EM2 | GIT epithelium EM3 | Desmosome EM Adhesion Cartoons: Close Intersection | Adherens junction | Desmosome | Gap junction blastocyst flap - zona pellucida lost, ZP has a place of sperm entry, and the whole of the P.K. is broken down by the secretion of the uterus and possibly the secretion of the blastula. Uterine glands - sections needed for blastocist molility and nutrition Links: MBOC Figure 21-69. Blastula Blastocyst flap Around the fifth day, a human blastocist hatches from the protective zone of pellucida. This ripple allows for increased growth, access to secretion of uterine nutrients and blastocyst adhesion to the uterus lining. Associated with this hatching process are a series of physical shrinkages. In blastocyst, the reductions after the formation of blastogel are repeated, and the frequency of brushes is higher during the wave season than in the periods before and after hatching. [20] Interestingly, the same researchers in this mouse study suggest that weaker contractions (less than a 20% decrease in volume) as opposed to strong brushes (20% or more of volume reduction) have the opposite effect of inhibition. In the mouse model, the defined areas of the zone pellucida shedding different (24% mural site, 24% internal cell mass site, 17% equatorial site, and 35% of other sites). [21] Links: Blastocyst Day 5-6 Movie &lt;mediaplayer width=500' height=450' image=amp;g;File:Human\_blastocyst\_day\_5-6.mp4&lt;mediaplayer&gt; Human blastocyst contractions (day 5-6)[10] Molecule factors TEA DNA-domain binding. These factors are binding on the TEA/ATTS cognate binding site[22] TEF-3 - renamed u Tead1, I Tead4 Tead3 - expressed u syncytiotrophoblasts E-cadherin - Calcium ion dependent cell adhesion molecule, cell membrane adhesive proteins needed for morula compaction epithin - A type II transmembrane serine prosthesis, found in the mouse to reproduce murine during embryo preimplantation. Expressed from the 8-cell phase in contacts with blastomer and co-localism in moruli with E-kadherin. [23] Na, K-adenosine triphosphatase - Sodium potassium pump, which generates osmotic gradient for fluid flow u blastokoeel Zonula occludens-1. (ZO-1) Close suspent proteins u moruli to translocista Transformation u mouse [24] Blastocist u Other species Mouse Blastocist Early gene expression[25] Early gene expression[25] Early gene expression[25] Early gene expression[25] Early mouse development model[26] Links: Mouse Development Bovine Blastocist KRT18 , FN1 i MYL6 term Bovine blastocist KRT18 i MYL6 term Linkovi. Cattle Development Reference i 1.0 1.1 Zhang P, Zucchelli M, Bruce S, Hamblili F, Stavrus-Evers A, Levkov L, Skottman H, Kerkela E, Kere J &amp;amp; Hovatta O. (2009). transcription profiling of development prior to human implantation. PLOS 1, 4, e7844. PMID: 19924284 DOI. ^ Sasaki H. (2015). The position- and polarity-dependent Hippo signaling regulates cellular destiny in the pre-implantation of mouse embryos. Semin. Cell Dev. Biol. 47-48, 80-7PMID: 25986053 DOI. ^ Rayon T, Menchero S, Nieto A, Xenopoulos P, Crespo M, Cockburn K, Cañon S, Sasaki H, Hadjantonakis AK, de la Pompa JL, Rossant J &amp; Manzanares M. 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