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enzyme and cystathionine synthase were present. None of these enzymes of transsulfuration was present in the placenta. Thus cyst(e)ine may be an essential amino acid in the immature human until sometime after birth. A single full term infant who died at 7 hr had 7% of normal cystathionase activity in the liver. Amino acid analyses of simultaneously obtained maternal and fetal blood, amniotic fluid, placenta, fetal liver, and fetal brain were performed. No consistent trends were noted during the period studied. However, the fetal-maternal ratio of all plasma amino acid concentrations at time of abortion was high (as at term). The highest ratio, by far, was that of cystine. In some cases, cystine was not measurable in the blood of the mother. Furthermore, in contrast to the mature human, cystathionine, the substrate for cystathionase, was higher in fetal liver than in fetal brain. In spite of this accumulation of cystathionine in fetal liver, there was none measurable in fetal or maternal blood nor in amniotic fluid. These studies suggest that the human fetus is entirely dependent on a maternal source of cvst(e)ine and that prematures and perhaps even full term newly born infants are dependent upon dietary sources of cyst(e)ine. Human milk is a high cyst(e)ine, low protein formula, whereas cow's milk is a low cyst(e)ine, high protein formula. Thus, these studies suggest that premature infants fed a high protein cow's milk formula retain more nitrogen and grow faster than infants fed a cow's milk formula containing lower amounts of protein, closer to that found in human milk, because the amount of cyst(e)ine, rather than total nitrogen, may be a limiting factor for protein synthesis. These results may also afford an enzymatic explanation for the transient hypermethioninemia seen in infants on high protein diets.

20. a₁-Fetoprotein, an index of maturation? R. LARDINOIS, D. ANAGNOSTAKIS, and M. ORTIZ. Gentre de Recherches Biologiques Neonatales, Paris, France.

It is known that in the human conceptus, serum a₁-fetoprotein reaches a maximal concentration at approximately 13 weeks, then decreases and disappears 1 or 2 weeks after birth. In the first step of this work, the existence of a₁-fetoprotein has been studied in the serum of three groups of neonates (premature, full term, and small for date). Electrophoresis on polyacrylamide gels and immunoelectrophoresis with a specific antibody against human a_ifetoprotein have been chosen: since they have a different degree of sensitivity, they can be used as a semiquantitative test. At birth, small for date infants have no a₁-fetoprotein or a very low concentration; the concentration is higher in full term babies and still higher in prematures. These observations suggest that a distinction between small for date and premature babies is possible by such a procedure. In the second step of this study, Mancini's immunochemical method for a₁-fetoprotein quantitative estimation is in process in order to see, especially in premature babies, whether serum a₁-fetoprotein concentration at birth is directly related to the length of gestation. If so, we shall have a simple biological test for the assessment of the gestational age of newborn infants.

21. Effects of varying severity of growth retardation on organ weight and cell population in fetal rats. J. S. Wigglesworth. *Hammersmith Hosp., London, England*.

Recent experimental studies have shown that a growth-retarding stress applied early in life results in diminished cell populations of all organs whereas a similar stress applied later on causes reduction predominately in cell size. The object of the present study was to determine the effects on cell size and population of organs of varying the severity of stress at a single time

interval in pregnancy. Unilateral uterine ischemia was induced surgically in rats on the 17th day of pregnancy to produce a range of fetuses with birth weight reduced by up to 45% of the values for fetuses from the control uterine horns. Cell size and populations at term were estimated from the figures for organ weight and total DNA. A close correlation was shown between the severity of growth retardation and the degree of reduction in organ weight and cell population although the size of the effect varied for different organs. Reduction in fetal weight by 45% reduced liver weight by 60% and liver cell population by 50%, whereas brain weight was reduced by only 20% with a 12% reduction in cell population. No consistent change was seen in weight or cell population of the placenta. It is concluded that for a growth-retarding stress acting at a single time interval in pregnancy the reduction in cell size and populations of different internal organs is directly related to the reduction in birth weight.

22. 1gA deficiency—hereditary aspects, P. Pelkonen and E. Savilaitti, Children's Hosp., Univ. of Helsinki, Finland.

Familial cases of selective immunoglobulin A deficiency have been documented, but the mode of inheritance remains unsettled. The propositus of the present study, a 9-year-old girl with a past history of frequent upper respiratory tract infections, had an unusually low serum IgA level, between 0.5 and 2.0 mg/100 ml (i.e., IgA detectable by double diffusion but not by the radial immunodiffusion technique). The mother of the patient showed a total lack of serum IgA and suffered from chronic urinary tract infection. The father and brother of the propositus and the maternal relatives tested had normal IgA levels. Both patients had normal karyotypes. In the propositus, IgA was detected in saliva and intestinal juice, but IgM was present in higher concentrations in these secretions. Direct immunofluorescent studies of both rectal and small intestinal mucosa revealed IgA-containing cells, but IgM-containing cells were predominant. In the mother, no IgA was detectable in whole saliva concentrated 20 times, and the rectal mucosa was completely devoid of IgA-containing cells, whereas IgM-containing cells were abundant. These findings suggest that IgA deficiency may differ in degree of severity. The daughter in this family could be heterozygous for a recessive trait, while the mother is homozygous. On the other hand, an autosomal dominant mode of transmission with variable gene expressivity cannot be ruled out.

23. Penicillin and dinitrophenyl antibodies in newborns and mothers detected with chemically modified bacteriophage. S. Levin, Y. Aleman, and M. Sela, Kaplan Hosp, and The Weizman Inst. of Sci., Rehovot, Israel.

The development of newer, highly sensitive techniques for the detection of small amounts of antibodies opens new fields for investigation. The immunospecific inactivation of chemically modified bacteriophages by antibodies directed toward the attached hapten or protein allows for the detection of as little as 0.2–2.0 ng antibody/ml. Likewise, minute amounts of hapten or protein may be detected and measured by their ability to inhibit the inactivation of the chemically modified bacteriophage by the antihapten or antiprotein antibodies. We have studied the presence of penicillin and DNP antibodies in the sera of newborn infants and their mothers by the use of penicilloylated and dinitrophenylated-T4-bacteriophage. In almost every paired sera studied, evidence was found for the presence of penicillin antibodies. Premixing the sera with penicillin eliminated the penicillin-T4-phage inactivation. Preheating the diluted sera for