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DEMONSTRATION OF CIRCULATING IMMUNE COMPLEXES (CICs) CONTAINING HUMAN THYROGLOBULIN (HuTg) IN A PATIENT WITH IMMUNE COMPLEX GLOMERULONEPHRITIS (ICGN)

MEDIATED BY THYROGLOBULIN ANTI-THYROGLOBULIN IMMUNE COMPLEXES. Jordan, S.C., Buckingham, B.A., Olson, D.L., Perelman, A.J., Kogut, M.D., and Fine, R.N., Univ. So. Calif. Sch. Med. and Childrens Hospital of Los Angeles, California

A pediatric patient (pt) with Graves disease and circulating anti-Tg antibody (ab) levels developed ICGN characterized by mesangial, subendothelial and subepithelial dense deposits by electron microscopy. Renal cortex tissue examined by direct immunofluorescence (IF) demonstrated granular glomerular immunoglobulin (Ig) and complement (C') deposition. Indirect IF demonstrated granular glomerular and mesangial deposits of HuTg. Indirect IF also showed that eluates of the pt's kidney contained (ab) reactive with thyroid follicular cells and colloid. CICs were demonstrated in the pt's sera by both the Raji cell assay and Clq solid phase assay. The indirect IF Raji assay demonstrated HuTg in ICs bound to the Raji cells. That HuTg was the antigenic component of CICs was further documented by a marked decrease Raji cell binding of CICs in the pt's sera when the sera was preincubated with HuTg. ICs eluted from both renal cortex tissue and CICs bound to Raji cells were analyzed by immunoelectrophoresis techniques and found to contain HuTg. We concluded that HuTg-anti-HuTg CICs have nephritogenic capacity and are important in induction and perpetuation of ICGN.

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4 of 6 patients with Alport syndrome from 3 different families showed presence of antithyroid antibody (ATA), and absence of anti DNA and anti nuclear antibody. 4 healthy members of these families, 22 patients with familial nephritis without deafness from 7 families and 10 patients with sporadic hematuria showed negative ATA. In general adult population 16 of 1000 subjects have positive titer of ATA. When often reported data are taken into account together with this survey, 14 of 19 patients with Alport syndrome have positive titer of ATA, indicating significant higher incidence of ATA in the disease.

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IMMUNOLOGIC STUDIES IN SHUNT NEPHRITIS CAUSED BY MICROCOCCUS AND STAPH. ALBUS INFECTION. Leumann E.P., Briner J., Joller P., Odermatt B., Seger R. Dept. of Pediatrics and Pathology, University of Zurich, Switzerland.

Two pediatric patients with shunt nephritis, one infected by Micrococcus (M), and the other by Staph. albus (S), were studied. Both exhibited cryoglobulinemia, elevated IgG indicating chronic infection, and very high agglutinating antibody titers (1:4000) against the respective organisms (M and S) cultured from the patients' blood. After revision of the shunts, the half life of the specific antibodies was determined.

Renal biopsies disclosed membranoproliferative glomerulonephritis type I (M) and type III (S). Granular deposits containing Clq, C3, IgM and IgG were detected at immunofluorescence examination. The glomerular immune deposits are currently being investigated for the presence of micrococcal and staphylococcal antigens.

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A BIOLOGICAL ACTIVITY OF IMMUNE COMPLEXES DEPOSITED IN GLOMERULI OF VARIOUS RENAL DISEASES IN CHILDHOOD. Takada, T., Yamagihara, T., Yamamoto, T., Morita, T., Kihara, I., Dept. of Pediat., Yoshida Prefectural Hospital, and Dept. of Pathol. Inst. Nephrol. Miigata Univ. Sch. Med., Miigata Japan.

In order to determine the biological activity of IC in glomeruli, we studied about 40 renal patients with the method Yamamoto described. 40 cases diagnosed by renal biopsy were IgA nephropathy(3), LN(5), Henoch-Schoenlein nephropathy(9), MPGN(7), MN(2), AGN(5), and lipid nephrosis(4). Among 40 patients kidney biopsies were obtained at intervals from 21 cases. A biological activity of IC was demonstrated with the number of polymorphonuclear leukocytes(PMN) attached in patient's glomeruli. In patients with lipid nephrosis, attachment of PMN was not seen. In MPGN, purpura nephritis and LN, the number increased significantly. Though, among the immunofluorescent findings of kidney biopsies obtained at intervals from 21 cases, 18 cases were almost similar, the number of PMN attached in glomeruli decreased in the cases with the clinical improvement. These results indicate that this method is useful to observe the degree of histological and clinical activity of glomerulonephritis in re-biopsies.

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CIRCULATING IMMUNE COMPLEXES(CIC) IN ACUTE POSTSTREPTOCOCCAL GLOMERULONEPHRITIS(APSGN) AND STEROID SENSITIVE NEPHROTIC SYNDROME(SSNS): Popović-Rolović, M., Živanović, Lj., Nikolić, V., Milošević-Jovčić, N., Children's University Hospital and Institute for Medical Research, Beograd, Yugoslavia

The purpose of this study is to report the findings of CIC using Polyethylene glycol precipitation technique(PEG) and modified variant of inhibition of RF Latex agglutination test(RF-inh.) in 18 patients with APSGN, in 18 postnephritic children who recovered from the acute phase of APSGN 2-4 years earlier and 18 patients with SSNS.

CIC were found in 14/18 patients with APSGN when tested during the first three weeks of the disease using PEG and in 11/13 using RF-inh. Sequential determination of CIC showed the highest values of CIC within 3-5 weeks of the disease. In acute phase gross hematuria was found in 13/14 patients with CIC and only in 1/4 without CIC; there was no difference in severity of proteinuria and serum creatinine. In 18 postnephritic children CIC were not found. In patients with SSNS, CIC were found in 12/14 while relapsing. During the follow-up these 12 patients had 26 relapses and CIC were found in 17/26 and 10/21 using PEG and RF-inh. respectively. In remission CIC were found only occasionally.

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DETECTION OF CIRCULATING IMMUNE COMPLEXES IN PATIENTS WITH GLOMERULONEPHRITIS

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Circulating immune complexes(CIC) are involved in the pathogenesis of various renal diseases. In eighty children with renal disease, CIC were measured using the Raji cell immunofluorescent method(Theofilopoulos's method). This method gave us high positive results in patients with acute glomerulonephritis(AGN) and chronic glomerulonephritis(CG). The type of CIC antibodies were checked again employing precipitation tests using the polyethylene glycol(PEG) technique(Abe's method). The major results are summarized as follows: 1) In AGN, high levels of CIC were detected especially within 4 weeks after the onset of the disease. CIC including IgA, as well as, IgG and IgM were often detected using the PEG technique, but the level of CIC including IgA were higher than the others. 2) The levels of CIC in poststreptococcal infection, without renal involvement, were intermediate-level between AGN and healthy children. 3) In nephrotic syndrome, except in one case with positive HBs antigen, CIC were not detected even in acute phase. 4) In CG, CIC were detected in high levels, and there was a correlation, the pattern of glomerular localization of immune complexes and CIC. The observations using Raji cell method and PEG technique may be a useful method to monitor activities of disease in patient with immunologically mediated renal diseases.