
studies in the pathogenesis of glomerulonephritis

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Saprophytic and non-pathogenic streptococci are widely distributed in nature, but some species are pathogens for human being, among of which are, group A, beta hemolytic streptococci which are common in sore throat of children. This infection is usually of short duration but it always produce delayed acute post streptococcal glomerulonephritis. Acute post-streptococcal glomerulonephritis (AGN) is a form of acute glomerular injury which is a delayed sequela of infection with certain nephritogenic strains of group A beta hemolytic streptococci. Documentation of streptococcal infection (positive throat culture or elevated titres of antibody to certain extracellular antigens such as streptolysin O) is an important factor for establishing the diagnosis. The urinalysis usually reveals proteinuria, hematuria, and the presence of red blood cell casts. Hypertension, azotemia, and oliguria are usually mild but may be severe in a small percentage of patients. AGN is generally a nonprogressive disease in children leading to complete recovery (Baldwin et al., 1974). Many morphological, clinical, and serological features suggest that acute poststreptococcal glomerulonephritis (APSGN) is an immune complex disease (Glassock et al., 1966). Frequently following the onset of this disease, serum C3 levels are markedly depressed. Discrete deposits of IgG and C3 on the epithelial side of the GBM have been detected. In addition, soluble immune complexes have been detected in the circulation of some patients with AGN (Mohammed et al., 1977; Ooi et al., 1977). Up till now, no specific genetic marker is characteristic for post-streptococcal glomerulonephritis in Egypt. As striking relationships have been found between HLA-antigen and many diseases, the present work was undertaken to look for any association between (PAGN) and certain alleles of the HLA-A, B, C and DR. The present work included 33 patients, 19 males and 14 females with age ranged from 4-14 years. They were taken up consecutively from the inpatients of the Department of Pediatrics at Mansoura University Hospital. The cases were presenting clinically and laboratory as acute glomerulonephritis, post-streptococcal on the basis of presence and/or history of sore throat, or pyogenic skin infection, and evidence of streptococcal infection. For all members included in the study, the following were done:- Full detailed history. Laboratory investigations . • Culture of skin infection and throat swabs, and serological grouping of isolated streptococci by: 1) Streptex test 2) Bacitracin sensitivity test. * Urine analysis_ Chemical and serological investigations. * serum creatinin, creatinin clearance serum electrolyte to evaluate kidney function. * C-reactive protein and A.S.O titre. * Identification of serum (Igs), complement and CIC. * HLA tissue typing. In the present study, throat swabs were taken from 33 cases of acute glomerulonephritis. Beta-hemolytic streptococci were

identified by the presence of beta-hemolysis on blood agar plates. The number of beta-hemolytic streptococcal isolates was 7 in all 33 cases. These isolates were grouped by streptex kits, bacitracin sensitivity disks. The number of group A beta-hemolytic streptococci identified by streptex was 7 (100%) of the isolated cases. The number of group A identified by bacitracin disks was 6 out of 7 seven cases (85.7%). From the previous data, it is quite clear that the incidence of group-A beta-hemolytic streptococcal isolates is low. The sensitivity of streptex kits in identification of group A streptococci was striking, but bacitracin disks were less sensitive. The antistreptolysin O titre was determined by the latex method. The mean value of antistreptolysin O titre was higher in cases from which beta-hemolytic streptococci were isolated than in cases from which no beta-hemolytic streptococci were isolated. The mean value of antistreptolysin O titre was (742.86 & 384.62) respectively. C-reactive protein was found to be positive in all cases of acute poststreptococcal glomerulonephritis. HLA antigens were typed by using the microlymphocytotoxicity tissue typing technique (Bodmer, 1978). The HLA B8 is the only antigen in locus B that shows significantly higher frequency compared to controls because its frequency in the patient (42%) compared to normal controls (6.5%) is significantly high and $P < 0.001$. Also the relative risk is high and significant ($RR = 10.6$). Further, the contribution of the antigen as measured by etiologic fraction is high 0.684. Statistical analysis also revealed that the only significant difference of HLA-DR is that between the frequencies of HLA DR3. The DR3-antigen frequency in the patients (27.3%) compared to that of controls (14.9). The $RR = 7.14$. Further, the test of significance of the RR (total X^2) shows significance RR . $P < 0.001$. Assays for circulating immune complexes by laser nephelometry were positive and more than 1.5 mg/dL in every patient which indicated antigen-antibody reaction forming the (222) immune complexes in all patients. The level of C3 measured by laser nephelometry was found decreased in 23 out of 33 cases. The means of C3 and C4 levels were consequently the following, 51.838 and 32.65. C3 was present with seven cases having HLA-B8 and decreased in 6 out of these 7 cases while was decreased in 4 out of 6 cases having HLA-DR3 haplotype, and was decreased in all the six cases having HLA B8/DR3. This indicated that there is a great association of both C3 and HLA B8 as well as B8/DR3 haplotypes in APSGN. Serum levels of IgG were below the normal level in 28 out of 33 cases. IgG was decreased in seven cases out of eight, having HLA-B8 and with 5 out of 6 having HLA-DR3. This indicates that there is a great association of the IgG and HLA B8 as well as HLA-DR3 in APSGN. In contrast, the IgA, was decreased in only one case out of six with B8, two cases out of six with DR3 and three cases out of six with B8 DR3. This indicated that IgA is not associated with any genetic factor in APSGN. Also there is insignificant correlation between the levels of IgM and HLA haplotypes, however the mean of IgM in our patient (301.3) was high than that of control (164.2).