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□□□□□□□□□□ Hong Kong, 2012-10-26 CD-ROMs in Print ,2003 **Index Medicus** ,2003 Vols for 1963 include as pt 2 of the Jan issue Medical subject headings Cross-Reactive Immunity and COVID-19 Alberto Beretta, Ahmed Yaqinuddin, Aristo Vojdani, Pedro A. Reche, 2025-01-06 There is a growing evidence that cross reactive immunity from common human coronaviruses hCoVs can shape the immunological response to SARS CoV 2 Cross reactive T cells appear to play a protective role against disease but there is no consensus on the role of cross reactive antibodies with some authors reporting protective effects mechanisms and other authors reporting evidence of negative interference due to the generation of low affinity antibodies in the context of an original antigenic sin Reports of association of pre existing immunity to hCoVs and milder clinical manifestations of COVID 19 conflict with other reports showing in COVID 19 patients a strong back boosting of antibodies to hCoVs with very little if no neutralizing activity against SARS CoV 2 However SARS CoV 2 hCoVs cross reactive antibodies more specifically those directed against the S2 subunit of the Spike protein may be protective via Fc mediated mechanisms that recruits NK cells ADCC or similar mechanisms Since these antibodies do not neutralize or neutralize very poorly SARS CoV 2 in vitro they have been only occasionally investigated and rarely considered in the search for correlates of protection Alternatively it should also be considered that protection against severe COVID 19 disease may be sustained by cross reactive T cells with cross reactive antibodies being only a proxy of protective T cell immunity Although pre existing immunity to SARS CoV 2 is more often linked hCoVs other sources of cross reactive immunity to SARS CoV 2 including bacteria and vaccine antigens are also plausible and should be considered In particular it has been identified first computationally and later experimentally cross reactive immunity to SARS CoV 2 from combination tetanus diptheria vaccines DTaP DTwP TD and Tdap Moreover epidemiological studies have clearly shown that subjects vaccinated with tetanus diptheria vaccines are less likely to suffer severe COVID 19 Cross reactive immunity to SARS CoV 2 from pediatric tetanus Diptheria vaccines and seasonal exposures to hCoVs likely play an important role in protecting children from COVID 19 On the other hand the elderly may fail to mount an effective primary response to SARS CoV 2 following infection owing to the loss of cross reactive memory B and T cells and their relative inability to recruit naive T and B cells **Byte** ,1998

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