

## Sample of Level 2 English Editing

Field of research: Medicine

### High avidity cytokine autoantibodies in health and disease: pathogenesis and mechanisms

#### INTRODUCTION

In 1981, antibodies against interferon (IFN)- $\alpha$  or IFN- $\beta$  were reported in participants of ~~—~~ clinical trials evaluating administration of these exogenous recombinant human cytokines<sup>1,2</sup>. Less than a decade later, naturally occurring autoantibodies against endogenous interleukin-1 $\alpha$  (IL-1 $\alpha$ ) were reported<sup>3,4</sup>. Cytokine autoantibodies have been identified in pharmaceutical preparations of intravenous immunoglobulin (IVIG) and in the healthy subjects from ~~which-whom~~ IVIG ~~is-were derived~~ prepared<sup>5-9</sup>. Numerous studies have additionally demonstrated that cytokine autoantibodies ~~can-are~~ detectable ~~be-detected~~ in health and disease in various tissues, including blood,<sup>3,4,7,10-14</sup> lung,<sup>10,15-17</sup> pleural fluid,<sup>18</sup> central nervous system,<sup>19</sup> gums,<sup>20</sup> and synovial fluid.<sup>21,22</sup> In some cases, specific autoantibodies have been implicated in ~~or-predisposing-to~~ disease pathogenesis or predisposition.<sup>10,14,16-18,23-27</sup> Others appear to ~~cause~~ be responsible for the ~~the~~ associated physical manifestations of disease.<sup>28-30</sup> In addition to blocking ~~the~~ specific cytokine function, cytokine autoantibodies may play important roles in health and disease by virtue of their ability to form immune complexes<sup>12,13,16,17,31-33</sup>, which can interfere with their detection by conventional cytokine 'antigen-capture' methods<sup>12,13</sup>. The rRecent development of novel methods has improved our ability to detect cytokine autoantibodies and ~~therefore-has enhanced~~ our understanding of their ~~real-and-potential~~ role and functions ~~in-health-and-disease~~. Here In this report, we reviewed ed the current literature ~~documenting-relating to~~ the presence, biological function, potential mechanisms of production and significance of cytokine autoantibodies.

**Comment [SU1]:** CHECK: It would be a good idea to use a reference formatting program such as Endnote to ensure your references are all formatted the same. You have some with the references after the full stop and some with them before the full stop. Consider checking through your in-text citations in order to maintain consistency in reference formatting.

**Comment [BeK2]:** CHECK: I assume by manifestations you are referring to the physical (clinical) manifestations of the disease. I have clarified this for you in the edit.

# CYTOKINE AUTOANTIBODIES IN DISEASE

Cytokine autoantibodies have been ~~reported~~ detected in patients with in various diseases. (Table 1). ~~To better understand~~ To explore the pathogenic roles of cytokine autoantibodies in each of these diseases, we have discussed below ~~described~~ (1) introduction ~~the~~ role of a particular ~~each~~ the cytokine and in the given disease, (2) the frequencies ~~incidence of detection~~ of ~~the~~ autoantibody yields ~~detection~~ in the disease, (3) reports ~~identifying a~~ the mechanism of their function in disease pathogenesis, and (4) reports dealing with the association of ~~the~~ autoantibodies with certain therapies or clinical manifestations. In each case, the effects on disease pathogenesis or severity are ~~were~~ linked to the biologic function(s) regulated by the specific cytokine and the interrelationship of this regulation with other members of the cytokine regulatory network (Figure 1).

In this section, the actions of cytokine autoantibodies associated with diseases are ~~were~~ classified into four categories. Firstly ~~The first~~ category describes, ~~the~~ autoantibodies highly ~~that effectively~~ abrogate ing cytokine activity and are largely ~~can be~~ the primary cause or predisposing factor of disease pathogenesis, when they are ~~more abundant than~~ present in levels over a certain critical threshold,<sup>12</sup> ~~or can be predisposing to the diseases.~~ (Figure 2A). Secondly ~~Autoantibodies of the second category involved those,~~ increases in some cytokine autoantibodies ~~which~~ exacerbate certain diseases by augmenting cytokine activity ies (Figure 2B). The third category consisted of ~~Thirdly,~~ those cytokine autoantibodies that may attenuate the severity of certain diseases. (Figure 2C), while the last category. Finally described those whose, ~~although the~~ roles of cytokine autoantibodies are unknown, although their induction is thought likely to be triggered by certain diseases.