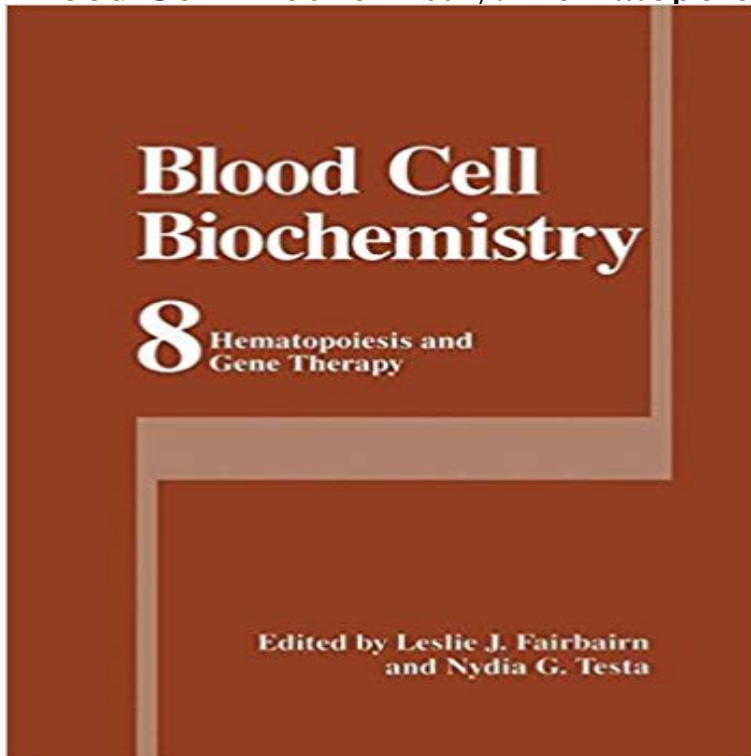


Blood Cell Biochemistry: Hematopoiesis and Gene Therapy



Since the first concepts of gene therapy were formulated, the hemopoietic system has been considered the most natural first target tissue for genetic manipulation. The reasons for this include the fact that a very large number of inherited disorders (including some of the most common disorders, such as the hemoglobinopathies) are disorders of the hemopoietic system, and the large amount of experience in hemopoietic transplantation biology. The consequence of this resulted in the first clinical trial of gene therapy in 1989, where two children suffering from severe combined immune deficiency (ADA-SCID) were transplanted with T-cells expressing adenosine deaminase (the defective enzyme in patients with this disorder). The partial success of this treatment was perhaps responsible for undue optimism among those proposing other gene therapy treatments within the hemopoietic system, and it has since become clear that there are a number of technical and biological difficulties to overcome before hemopoietic gene therapy becomes a mainstream therapeutic strategy. The chapters in this book evaluate the need for gene therapy in the hemopoietic system, discuss how efficient gene transfer and expression can be achieved in the target cells, highlight areas of difficulty to be addressed, and examine a number of potential applications of the gene therapy approach. The book begins with a chapter by Testa and colleagues, discussing the various sources of hemopoietic cells for both transplantation and gene therapy.

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