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Antibody Engineering Volume 1

Edited by Roland Kontermann and Stefan Dübel Second Edition



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Sudhir Paul

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exquisite binding specificity of antibodies has made them valuable tools from the laboratory to the clinic Since the description of the murine hybridoma technology by K hler and Milstein in 1975 a phenomenal number of mo clonal antibodies have been generated against a diverse array of targets Some of these have become indispensable reagents in biomedical research while others were developed for novel therapeutic applications. The attractiveness of an bodies in this regard is obvious high target specificity adaptability to a wide range of disease states and the potential ability to direct the host s immune s tem for a therapeutic response The initial excitement in finding Paul Ehrlich's magic bullet however was met with widespread disappointment when it was demonstrated that murine antibodies frequently elicit the human anti murine an body HAMA response thus rendering them ineffective and potentially unsafe in humans Despite this setback advances in recombinant DNA techniques over the last 15 20 years have empowered the engineering of recombinant antibodies with desired characteristics including properties to avoid HAMA The ability to p duce bulk quantities of recombinant proteins from bacterial fermentation also fueled the design of numerous creative antibody constructs To date the United States Food and Drug Administration has approved more than 10 recombinant antibodies for human use and hundreds more are in the development pipeline The recent explosion in genomic and proteomic information appears ready to deliver many more disease targets amenable to antibody based therapy **Protein Misfolding and Disease** Peter Bross, Niels Gregersen, 2008-02-02 For decades it has been known that structured conformations are important for the proper functioning of most cellular proteins However appreciation that protein folding to the functional conformations as well as the structural maintenance of protein molecules are very complex processes has only emerged during the last ten years. The intimate interplay uncovered by this scientific development led us to realize that perturbations of the protein folding process and disturbances of conformational maintenance are major disease mechanisms. This development has given rise to the concept of conformational diseases and the broader signature of protein folding diseases comprising diseases in which mutations or environmental stresses may result in a partial misfolding that leads then to alternative conformations capable of disturbing cellular processes This may happen by self association aggregation as in prion and Alzheimer's diseases or by incorporation of alternatively folded subunits into structural entities as in collagen diseases Another possibility is that folding to the native structure is impaired or abolished resulting in decreased stea state levels of the correctly folded protein as is observed in cystic fibrosis and 1 antitrypsin deficiency as well as in many enzyme deficiencies In addition deficiencies of proteins that are engaged in assisting and supervising protein folding protein quality control may impair the folding of many other proteins resulting in pathological phenotypes Examples of this are the spastic paraplegia attributable to mutations in mitochondrial protease chaperone complexes **Antibody Engineering** Damien Nevoltris, Patrick Chames, 2018-09-24 This detailed new edition provides complete and easy access to a variety of antibody engineering techniques. The volume explores topics such as the generation of native synthetic or immune antibody libraries the selection of lead candidates via the different powerful

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