1. Record Nr. UNINA9910298291003321 Autore De Arnab Titolo Ubiquitin Chains: Degradation and Beyond / / by Arnab De Pubbl/distr/stampa Cham:,: Springer International Publishing:,: Imprint: Springer,, 2015 3-319-14965-2 **ISBN** Edizione [1st ed. 2015.] Descrizione fisica 1 online resource (119 p.) Springer Theses, Recognizing Outstanding Ph.D. Research, , 2190-Collana 5053 574.19245 Disciplina Soggetti **Proteins** Cell cycle **Immunology Protein Science** Cell Cycle Analysis Lingua di pubblicazione Inglese **Formato** Materiale a stampa Livello bibliografico Monografia Note generali Description based upon print version of record. Nota di bibliografia Includes bibliographical references at the end of each chapters and index. Nota di contenuto 1 Immune homeostasis: Activation and Downregulation of NF-B -- 2 Regulation of NF-B signaling by regulatory ubiquitination: specific pathways and A20 -- 3 The deubiquitinase activity of A20 is dispensable for its role in NF-B signaling -- 4 Perspectives on regulatory ubiquitination. This thesis examines the evidence for regulatory ubiquitination by Sommario/riassunto focusing on A20. It provides an insightful and in-depth evaluation of the current literature by critically examining the evidence of K63-linked regulatory ubiquitination in regulating cell-signalling. It is also the first thesis to directly test the role of regulatory ubiquitination in NF-kB signaling in vivo. The case for regulatory ubiquitination has been to a large extent predicated upon the presumed deubiquitinase activity of A20, long considered a key regulator of inflammatory responses as mice lacking A20 die from multi-organ inflammation and cachexia. The theses reports the creation and characterization of a knock-in mouse that expresses a mutated form of A20 which selectively lacks the

deubiquitinase activity. The knock-in mice surprisingly display completely normal NF-B activation with no accompanying

inflammatory phenotype. Given that the presumed role of A20 as a deubiquitinase has been used to support the importance of regulatory K63-linked ubiquitination in NF-kB signaling, this study will help focus future research efforts into alternative target pathways that do not depend on K63 ubiquitination. In fact, the work suggests that it might be important to revisit the role of K63-linked polyubiquitination in cellsignalling. Ubiquitin Chains: Degradation and Beyond is essential reading for anyone conducting research in cell-signalling and immunology. Dr. Arnab De received his PhD from the Department of Microbiology & Immunology at Columbia University. During his PhD, he developed transgenic mice to study the mechanism of action of a critical tumor-suppressor called A20. He is also well known for having developed peptide-based prodrugs as therapeutics for diabetes. His work has been reported by the media, and has resulted in multiple patents and publications in peer reviewed journals. He presented his findings at the American Peptide Symposium and was awarded the Young Investigator's Award. He is the author of the book entitled Application of Peptide-Based Prodrug Chemistry in Drug Development, with a foreword written by Professor Jean Martinez (Former President, European Peptide Society) and published in the series SpringerBriefs in Pharmaceutical Science & Drug Development. His research interests lie at the intersection of chemistry and medicine. Besides biomedical research, he is also generally interested in public health policy and general scientific outreach.