Food Allergy Molecular and Clinical Practice Andreas L. Lopata (ed.)



A SCIENCE PUBLISHERS BOOK

Food Allergy Molecular and Clinical Practice

Editor

Andreas L. Lopata

James Cook University College of Public Health, Medical & Veterinary Sciences Centre of Biodiscovery and Molecular Development of Therapeutics Douglas, Queensland, Australia



CRC Press is an imprint of the Taylor & Francis Group, an informa business A SCIENCE PUBLISHERS BOOK Cover photograph reproduced by kind courtesy of Dr. Sandip Kamath.

CRC Press Taylor & Francis Group 6000 Broken Sound Parkway NW, Suite 300 Boca Raton, FL 33487-2742

• 2017 by Taylor & Francis Group, LLC CRC Press is an imprint of Taylor & Francis Group, an Informa business

No claim to original U.S. Government works

Printed on acid-free paper Version Date: 20170212

International Standard Book Number-13: 978-1-4987-2244-5 (Hardback)

This book contains information obtained from authentic and highly regarded sources. Reasonable efforts have been made to publish reliable data and information, but the author and publisher cannot assume responsibility for the validity of all materials or the consequences of their use. The authors and publishers have attempted to trace the copyright holders of all material reproduced in this publication and apologize to copyright holders if permission to publish in this form has not been obtained. If any copyright material has not been acknowledged please write and let us know so we may rectify in any future reprint.

Except as permitted under U.S. Copyright Law, no part of this book may be reprinted, reproduced, transmitted, or utilized in any form by any electronic, mechanical, or other means, now known or hereafter invented, including photocopying, microfilming, and recording, or in any information storage or retrieval system, without written permission from the publishers.

For permission to photocopy or use material electronically from this work, please access www.copyright.com (http://www.copyright.com/) or contact the Copyright Clearance Center, Inc. (CCC), 222 Rosewood Drive, Danvers, MA 01923, 978-750-8400. CCC is a not-for-profit organization that provides licenses and registration for a variety of users. For organizations that have been granted a photocopy license by the CCC, a separate system of payment has been arranged.

Trademark Notice: Product or corporate names may be trademarks or registered trademarks, and are used only for identification and explanation without intent to infringe.

Library of Congress Cataloging-in-Publication Data

Names: Lopata, Andreas Ludwig, editor. Title: Food allergy : molecular and clinical practice / editor, Andreas Ludwig Lopata. Other titles: Food allergy (Lopata) Description: Boca Raton, FL : CRC Press, 2017. | "A Science Publishers book." | Includes bibliographical references and index. Identifiers: LCCN 2017005276] ISBN 9781498722445 (hardback : alk. paper) | ISBN 9781498722452 (e-book) Subjects: | MESH: Food Hypersensitivity--therapy | Allergens--chemistry Classification: LCC RC596 | NLM WD 310 | DDC 616.97/5--dc23 LC record available at https://lccn.loc.gov/2017005276

Visit the Taylor & Francis Web site at http://www.taylorandfrancis.com

and the CRC Press Web site at http://www.crcpress.com

Preface

Allergy-related diseases are today recognized as reaching epidemic proportions, with up to 30% of the general population suffering from clinical symptoms ranging from urticaria, rhinitis and asthma to life-threatening anaphylactic reactions.

The main contributors to the increasing prevalence of allergy seem to be very diverse including increasing immunological predisposition ('atopy'), changing food consumption and well as living conditions. The dramatic increase of allergic diseases is not only seen in the developed world, but increasing evidence indicates that also developing countries are considerably affected. Already over fifty percent of the world population is living in Asia, where not only food consumption, but also food allergies are very different from what is mainly published from Western countries. In the research efforts in the field of food allergy two main questions are often asked: What makes one person allergic to a particular food and not the other? Furthermore, Why are some foods and food proteins more allergenic than others? In addition it is very difficult to predict the severity of clinical reaction and the amount of allergen required to elicit these reactions.

Major food allergens from a small number of sources were identified and purified as early as the 1970s. A boost in the number of newly identified allergens was elicited by the general availability of recombinant DNA technology in the late 1980s. The ever-growing IUIS Allergen Nomenclature Database contains currently over 840 allergens from 252 sources and their isoforms and variants. Currently we know about 290 food allergens from 98 different food sources. Recent developments into the molecular nature of allergenic proteins enabled us to classify most allergens into few protein families with limited biochemical function. Allergenic proteins can be classified into approximately 130 Pfam protein families, while the most important plant and animal food allergens can be found in 8 protein superfamilies and is discussed in detail in Chapters 1 and 2.

The correct diagnosis of a food allergy can be complex, but includes a convincing clinical history as well as the presence of elevated levels of specific IgE antibody to allergenic proteins in a given food. Therefore, detailed knowledge about the food specific allergenic proteins is central to a specific and sensitive diagnostic approach. The different allergens of peanut, egg, fish, shellfish and food contamination parasites and their diagnostic application are detailed in Chapters 3 to 7.

The food industry is one of the largest employers of workers with about 10% and therefore is the allergic sensitisation to food borne proteins at the workplace not surprising. Workers at increased risk of allergic sensitisation include farmers who grow and harvest crops; factory workers involved in food processing, storage and packing; as well as those involved in food preparation (chefs and waiters) and transport and is detailed in Chapter 8.

Research in food allergies and allergens is much more complex than investigating inhalant allergens since food proteins often undergo extensive modifications during food processing. Furthermore these allergenic proteins are embedded in a complex matrix and may undergo physicochemical changes during digestion and subsequent uptake by the gut mucosal barrier and presentation to the immune system, and have been highlighted in Chapter 9.

Furthermore, food processing results often in water-insoluble proteins, which makes the traditional serological analysis of allergenicity difficult as well as detection and quantification in the food matrix. The approaches and problems of quantifying allergen residues in processed food are detailed in Chapter 10.

To characterize allergens better but also develop better diagnostic and therapeutics, recombinant allergens are increasingly utilized. Unlike natural allergens or allergen extracts, the production of recombinant proteins is not dependent on biological source material composed of complex mixtures of allergen isoforms. The use of recombinant allergens has revolutionized diagnosis, enabling clinicians to identify disease eliciting allergens as well as crossreactivity pattern, thereby providing us with the tools necessary for personalized allergy medicine and therapeutics and is detailed in Chapter 11.

Food allergy is a growing problem globally carrying a huge socioeconomic burden for patients, families and the community. Although fatalities are fortunately rare, the fear of death is very real for each patient. Currently, there is no cure for any food allergy available, with management strategies focusing on complete avoidance and utilization of adrenaline as the emergency antidote for anaphylaxis. There is a very strong imperative for safe and effective specific therapeutics for food allergy and one strategy based on T-cell epitopes for peanut allergy is detailed in Chapter 12.

We hope that the joined effort by the authors will not only provide pragmatic information for current food allergy research but also serves as a foundation for significant new research that will advance our current knowledge.

Contents

Preface					
1.	Biomolecular and Clinical Aspects of Food Allergy <i>Heimo Breiteneder</i>				
	1.1	Introduction	2		
	1.2	 Prolamin Superfamily 1.2.1 Prolamins 1.2.2 Bifunctional Inhibitors 1.2.3 2S Albumins 1.2.4 Nonspecific Lipid Transfer Proteins (nsLTPs) 	3 4 4 5 6		
	1.3	Cupin Superfamily 1.3.1 Vicilins (7S globulins) 1.3.2 Legumins (11S globulins)	8 8 9		
	1.4	EF-hand Superfamily 1.4.1 Parvalbumins	10 10		
	1.5	Tropomyosin-like Superfamily	11		
	1.6	Profilin-like Superfamily			
	1.7	Bet v 1-like Superfamily			
	1.8	The Casein and the Casein Kappa Family			
	1.9	Calycin-like Superfamily 1.9.1 Lipocalins	15 16		
1	10	Conclusions	16		
1	Ackno	owledgement	17		
Ι	Refere	ences	18		
2.	Non Chris	nenclature of Food Allergens stian Radauer	30		
	2.1	Introduction	31		
	2.2 Allergen Nomenclature 2.2.1 Origin				

Food Allergy: Molecular and Clinical Practice

	2.2.2	Genus a	and Species Names	33
	2.2.3	Allerge	n Numbers	33
	2.2.4	Isoaller	gens and Variants	34
2.3	Subm Datab	itting Ne ase	w Allergens to the WHO/IUIS Allergen	35
	2.3.1	Allerge	n Source	36
	2.3.2	Sequence	ce Data	38
	2.3.3	Tested I	Patient Population	38
	2.3.4	Sensitiz	ation to the Submitted Allergen	38
2.4	Concl	usions		39
Refe	rences			39
Nut	Allerg	у		41
Dwa	n Price,	. Wesley B	Burks and Cenk Suphioglu	
3.1	Introc	luction		42
3.2	Why a	are Nut A	Allergens so Allergenic?	43
	3.2.1	Allerge	n Abundance	44
	3.2.2	Comple	ex Structural Integrity	44
	3.2.3	Special	Allergen Attributes	44
3.3	What	Therapie	es are Currently Addressing Nut Allergy?	45
3.4	Explo	ring Cau	ses of Nut Allergy	46
	3.4.1	Breakin	g Down Barriers	46
		3.4.1.1	Increased intestinal permeability	46
		3.4.1.2	Dermal barrier failure	49
	3.4.2	Initial A	Illergen Encounters—Is the Timing of	49
		Allerge	n Introduction Important?	
		3.4.2.1	In utero	49
		3.4.2.2	Breast milk	50
		3.4.2.3	Early foods	51
	3.4.3	Immun	e System Development—Preparing the	52
		Gut for	Nut Allergen Contact	
		3.4.3.1	The mucosal response to microbe	52
			colonization and gut development	
		3.4.3.2	Normal establishment of the	53
		0 4 0 0	microbiome	
		3.4.3.3	Living conditions	54
		3.4.3.4	birth type	54
		3.4.3.5	Infant feeding practices	55

3.

			3.4.3.6 3.4.3.7	Antibiotics Probiotics	55 57
	35	Concl	usions	110010400	58
	Refe	rences	abiono		50
	T	A 11			59
4.	Egg Paul	J. Turn	y er and Diı	anne E. Campbell	70
	4.1	Introc	luction		71
	4.2	Egg P	rotein All	ergens: Composition and Chemistry	71
		4.2.1 4.2.2	Egg Whi Egg Yolk	te <	72 74
	4.3	Prima	ry Preven	ntion of Egg Allergy	75
	4.4	IgE-m	ediated E	lgg Allergy	76
		4.4.1	Prevaler	ace and Natural History	76
		4.4.2	Diagnos	is	78
		4.4.3	Treatmen	nt	80
			4.4.3.1 4.4.3.2	Immunotherapy Vaccinations and medications containing Egg	80 81
	4.5	Non I	gE-Media	ted Food Allergy	82
		4.5.1	Food pro Syndrom	otein Induced Enteropathy ne (FPIES)	82
		4.5.2	Eosinopl	hilic Oesophagitis (EoE)	83
		4.5.3	Eczema		84
	Refer	rences			85
5.	Fish Anne	Allerg ette Kue	y hn and Ka	arthik Arumuqam	95
	5.1	Introd	luction	0	96
		5.1.1	Fish, a St	taple Food	96
		5.1.2	Adverse Allergy	Reactions to Fish: Intoxication and	97
		5.1.3	IgE-med	iated Fish Allergy: Clinical Phenotypes	100
		5.1.4	Fish Alle	ergy Diagnosis and Therapy	102
	5.2	Fish A	llergens		104
		5.2.1	Parvalbu	imins	104
		5.2.2	Fish Gela	atın	107

		5.2.3 5.2.4	Enolases and Aldolases Other Fish Allergens	108 109
	5.3	Transl 5.3.1 5.3.2	ational Aspects: From Bench to Bedside Allergen Contents in Food Fish Allergens	110 111 112
	Ackn	owledg	gement	114
	Refer	ences		114
6.	Rece Shell Sand	nt Adv lfish A ip D. Ka	vances in Diagnosis and Management of llergy amath, Roni Nugraha and Andreas L. Lopata	122
	6.1	Introd	uction	123
	6.2	Classi	fication of Shellfish	124
	6.3	Preval	lence of Shellfish Allergy	124
	6.4	Clinica	al Manifestations and Routes of Exposure	129
	6.56.66.76.86.9	Shellfi 6.5.1 6.5.2 6.5.3 6.5.4 6.5.5 6.5.6 6.5.7 Clinica Allerg Food I Conch	ish Allergens Tropomyosin Arginine kinase Myosin Light Chain Sarcoplasmic Calcium Binding Protein Troponin C Triose Phosphate Isomerase Paramyosin al and Immunological Cross Reactivity y Diagnosis and Management Processing and Effect on Allergens usions	132 133 134 134 134 134 135 135 135 135 140 143 145
	Ackn	owledg	gement	145
	Refer	ences		145
7.	Anis Fiona	a kis, A J. Bairt	llergy and the Globalization of Food d, Yasuyuki Morishima and Hiromu Sugiyama	155
	7.1	Introd	uction	156
	7.2	The Pa	arasite	157
	7.3	Anisal 7.3.1 7.3.2 7.3.3	kiasis: A Commonly Overlooked Infection Clinical Features Prevalence and Epidemiology Diagnosis and Treatment	158 158 160 162

		7.3.4	Allergy and Misdiagnosis of Fish Allergy Post-Infection	164
	7.4	Clinic of Foo	cal Implications of Travelling and Globalization od Products on Health	167
	7.5	Conc	lusions	169
	Refe	rences		170
8.	Occ Inha Moh	upation alant Fo amed F.	nal Allergy and Asthma Associated with ood Allergens Jeebhay and Berit Bang	176
	8.1	Introc Popul	luction—Food Industry and High Risk Working lations	177
	8.2	Food	Processing Activities and Allergen Sources	178
	8.3	Epide	emiology and Risk Factors	183
	8.4	Clinic	al Features and Diagnostic Approaches	189
	8.5	Biolog Occuj	gical and Biochemical Characteristics of known pational Allergens	191
		8.5.1	Seafood Allergens	191
		8.5.2	Flour Allergens Including Enzyme Additions	193
	0.6	0.3.3 D	spice Anergens	194
	8.6	Preve	ntive Approaches	194
	ð./	Conci	usion	196
	Kefe	rences		197
9.	The Food	Influe d Proce	nce of Dietary Protein Modification During essing on Food Allergy acek and Eva Untersmayr	203
	9.1	Introc	luction	204
	9.2	Food Diges	Protein Modification: From Processing to tion	206
	9.3	Thern	nal Food Processing	207
	9.4	Specif Allerg	fic Influence of Food Processing Methods on genic Food Compounds	209
		9.4.1	Peanut and Tree Nuts	209
		9.4.2	Milk	212
	~ -	9.4.3	Pollen Cross-reactive Food Allergens	213
	9.5	Chem Protei	ucal Food Modification: Nitration of Dietary	214

	9.6	Nitrati	on as a Concern in Food Allergy	216
	9.7	Furthe Oxidat	r Chemical Modifications: Reduction and tion of Food Proteins	217
	9.8	Conclu	isions	219
	Ackr	owledg	gements	220
	Refe	rences		220
10.	Dete and Sride	ection o Mass S evi Mura	f Food Allergen Residues by Immunoassays pectrometry alidharan, Yiqing Zhao, Steve L. Taylor and Nanju A. Lee	229
	10.1	Introd	uction	230
	10.2	Precau	tionary Labelling of Food Allergens	232
	10.3	Immu	noassays	234
		10.3.1	Enzyme-linked Immunosorbent Assay (ELISA)	234
		10.3.2	Non-competitive Assay for Food Analysis	235
		10.3.3	Competitive Inhibition ELISA	235
		10.3.4	Lateral Flow Devices (LFDs)	238
	10.4	Develo	opment of an ELISA	240
		10.4.1	Immunogen Preparation—Tree Nut Protein Extraction and Purification	240
		10.4.2	Antibody Production	243
	10.5	ELISA	Optimisation	246
		10.5.1	Coating and Blocking	246
		10.5.2	Buffer System, Incubation Time and Colour Development	247
		10.5.3	Cross-reactivity	248
		10.5.4	ELISA Validation	249
		10.5.5	Accuracy and Precision	249
		10.5.6	LOD, LOQ and Detection Range	250
		10.5./	Food Matrix Interference	250
	10 (10.5.8	Food Processing	251
	10.6	Mass S	pectrometry for Food Allergen Detection	255
		10.6.1	Clean-up	256
		10.6.2	Allergen Detection—Intact Proteins and Complex Mixtures	256
		10.6.3	Detection and Quantification of Allergen	258
			replices/ roteins in rood Using Mass Spectrometry	у

Contents

			10.6.3.1	Relative and absolute quantification	258	
			10632	Choosing suitable ionisation source	260	
			10.0.0.2	and mass analyser	200	
			10.6.3.3	Intensity and specificity of allergen	261	
				signatures		
			10.6.3.4	Synthetic peptides and isotopic	261	
				labelling		
		10.6.4	Food Al	lergen Signatures for Mass	262	
		1065	Spectro	metry Based Detection	266	
		10.0.5	Detectio	on rood Flocessing on rood Anergen	200	
		10.6.6	Protein	Glycosylation in Food Allergens	266	
		10.6.7	Multiple	exed Allergen Detection	268	
	10.7	Conclu	isions		270	
	Refe	rences			271	
11.	Reco Heid Fátin	ombinan i Hofer, 1 na Ferrei	n t Food A Anargyros ira, Gabrie	Ilergens for Diagnosis and Therapy <i>Roulias, Claudia Asam, Stephanie Eichhorn,</i> <i>le Gadermaier</i> and Michael Wallner	283	
	11.1	Introdi	uction		284	
	11.2	Recom	binant Fo	od Allergens	285	
	11.3	Physico Allerge	ochemical ens	Analysis of Recombinant Food	286	
	11.4	Immur Allerge	nological . ens	Analyses of Recombinant Food	327	
	11.5	Recom	binant Fo	od Allergens for Diagnosis	330	
		11.5.1	Peanut		332	
		11.5.2	Tree Nu	ts and Seeds	333	
		11.5.3	Fruits a	nd Vegetables	334	
		11.5.4	wheat		335	
		11.5.5	50y Fish		336	
		11.5.7	Shellfish	1	336	
	11.6	Recom	binant Fo	od Allergens for Allergy Therapy	337	
	11.7	Conclu	sions		341	
	Ackn	owledg	ements		341	
	Rofor	encee	Chiefito		340	
	References 34					

12.	Pear	ut Alle	rgy: Biomolecular Characterization for	351	
	Development of a Peanut T-Cell Epitope Peptide Therapy Jennifer M. Rolland, Sara R. Prickett and Robyn E. O'Hehir				
	12.1	Introdu	uction	352	
	12.2	Clinica	l Features of Peanut Allergy	353	
	12.3	3 The Mucosal Immune Response to Peanut Allergens			
	12.4	12.4 Allergenic Components of Peanut			
	12.5	Bioche	mical Properties of Peanut Allergens	357	
	12.6	Specifi	c Immunotherapy for Peanut Allergy	358	
	12.7	Develo 12.7.1 12.7.2	ppment of a SPIRE Therapy Rationale for SPIRE Therapy Validation of Allergen SPIRE Therapeutics in	359 359 361	
		12.7.3	Clinical Irials Mechanisms of Action of Allergen SPIRE Therapy	361	
	12.8	Design	n of a SPIRE Therapeutic for Peanut Allergy	363	
		12.8.1	Mapping T-cell Epitopes of Major Peanut Allergens	364	
		12.8.2	Determination of HLA-II Molecules which Present Peptides to T cells	365	
		12.8.3	Refinement of Peptides for Ease of Production and Solubility, Confirmation of T-Cell Reactivity and Lack of IgE-mediated Basophil Activation	365	
	12.9	Conclu	isions	366	
	Acknowledgements			367	
	References				
Ind	lex			373	