LONG-TERM FOLLOW UP OF INFANTS AT A HIGH RISK OF ASTHMA FROM A DEPRIVED COMMUNITY IN SOUTH WALES

January 2015

APPENDICES

Thesis submitted to Cardiff University for the Degree of Doctor of Medicine Dr Sadiyah Hand

Contents

Appendix page 1 Registrar General's Social Class based on occupation 2 2 Information capture form MAPS for birth and perinatal 3 period 3 Information capture form MAPS for examination at 3 6 months Information capture form MAPS for examination at 6 4 8 months 5 Information capture form MAPS for examination at 1 10 years of age Questionnaires on birthday cards, completed from 2nd 6 12 to 6th birthdays 7 Information capture for MAPS for examination at 7 13 years of age 8 Information capture form MAPS at 15 years of age 17 9 Information capture form MAPS for examination at 23 19 years of age 10 Core questions for asthma from ISAAC phase III 25 Skin prick testing method from ISAAC phase II manual 11 29-35

Appendix 1 Registrar General's Social Class based on occupation

Class Description Examples of occupations:

Non-manual

I Professional Doctors, chartered accountants, professionally qualified engineers

II Managerial & Managers, journalists, school teachers technical/intermediate

IIIN Skilled non-manual Clerks, cashiers, retail staff

Manual

IIIM Skilled manual Supervisors of manual workers, plumbers, electricians, goods vehicle drivers
 IV Partly skilled Warehousemen, security guards, machine tool operators, care assistants, waiters and waitresses
 V Unskilled Labourers, cleaners and messengers

Adapted from

http://www.statistics.gov.uk/downloads/theme_population/Life_Ex pect_Social_class_1972-05/life_expect_social_class.pdf Last checked 21st February 2010

FORM 2 : BASELINE INFORMATION

	s.,	ENTION STUDY	TYDFIL
Name	Hospita	1	Hosp.No.
Address			н.v.
1-7 	Reference number -Ist of twins; 2-2nd of Control - 2. More transporter INITIAL INTERVIEW Date of birth Date when first seen EDD 1 EDD definite; 2 EDD uncertain. FEEDING INTENSION Breast	$ \begin{array}{c} 32-33 \\ 34 \\ 35 \\ 35 \\ 35 \\ 34 \\ 34 \\ 34 \\ 34 \\ 36 \\ 1 \\ 2 \\ 3 \\ 4 \end{array} $	How many children has she had? <u>OCCUPATION</u> Husband's occupation; own occupation <u>SMOKING HABITS</u> Does mother smoke cigarettes? No if YES How many cigarettes per day? <5 5-14 15-24 25+ Did she stop smoking during pregnancy? Non-smoker Yes
2. 3. 28-30 217 29 Self. Hb. 0 0 1 1 2 2 3 3 4 4	Bottle Undecided HISTORY OF ALLERGY	<u>37-38</u> <u>39</u> 0 1 2	No If YES - gestation when she stopped <u>Type of Delivery</u> S.V.D. Breech Forceps
31 1. 2. 3.	 Asthma & eczema Eczema & hay fever Hay fever & asthma Asthma, eczema & hay fever MARITAL STATUS Single Married Widowed/Divorced	3	L.S.C.S. <u>Complications</u> No Yes

Forn 2 cont

Hospital no. Baby's name FEEDING ROUTINES COLS. Routine followed in hospital 60 41-50 Date of birth 0 Breast only 1 Breast & soya supplement Birth weight 2 Breast & CM supplement (kg x 100) Bottle - Soya Bottle - CM 3 4 SEX Specify CM ____ Male 1 51 2 Female Routine followed on discharge: CLINICAL EXAMINATION Breast only 61 0 Breast & Soya supplement Breast & CM supplement 1 0 Normal 2 52 Abnormal - specify Bottle - Soya Bottle - CM 1 3 4 Specify CM EXCLUSION FROM STUDY PERINATAL PERIOD Not excluded Stillbirth of Milconness 62 0 53 Normal 1 0 Baby unsuitable* Abnormal - specify 2 1 Mother unsuitable* 3 4 Early neonatal death 5 Non-cooperation 6 Other reason* Specify _ * 54-59 Date of discharge.

BIRTH, PERINATAL PERIOD AND DISCHARGE

4

ALLERGY PREVENTION STUDY : SUMMARY OF FEEDING AND ILLNESS

Baby's Name	Serial no.	1-5
	Card	2 6
Information checked with dietitian's card =	l; diary = 2, both = 3; neither = 4.	7

<u>5</u> .,	Mi 1k	feeding

Week	Milk (brand)	column	Week	Milk (brand)	column
1:			14		21
2 '		9	15		22
3.1		10	16		23
4		11	17		24
5.		12	18		25
6		13	19		26
7		14	20		27
8		15	21		28
9		16	22		29
10		17	23		30
11		18	24		31
12		19	25		32
13		20	26		33

Coding : Breast only = 0; Soya only = 1; Cow's milk only = 2;
Br. + soya = 3; Br. + CM = 4; Soya + CM = 5;
SMA liq. conc. only = 6; Br. + SMA Lc = 7; Other = 8; $NK = 9$.
Other combinations - code preferentially for (a) CM; (b) SMA 1.c.; (c) soya; (d) other.
First given - CM protein (any source exc. SMA LC) $34 35 36 37$ 38 39 40 41 42 43 Cereals $38 39eggs beef 1 other meat fish fish fish fruit/fruit juice (code week; not as yet = 99)$
First immunised for pertussis 5657 5859 6061 5253 5455 557
First suffered from cold cough cough ind charter of the state of the s
Treatment by a doctor for : respiratory condition (incl. cold, sore throat, otitis, cough) gastrointestinal condition other condition (viz. indult nytalis/typical or ophildusic cachii here) . Coding none o Vanhbichi = 1 - other treatment = ? . Coding none o Vanhbichi = 1 - other treatment = ? . Coding none of Nanhbichi = 1 - other treatment = ? . Coding none of Nanhbichi = 1 - ot

Appendix 3a

Form 6A

Baby's nameSerial No.Image: CardSerial No.Image: CardImage: Car		ALLERGY PREVE	NTION STUD	Y : E	EXAMINATION AT THREE MONTHS
13-20Veight (Kg x 100)Length (cm)Image: colspan="2">Code as follows :Length (cm)Image: colspan="2">Normal skin = 0Head circumference (cm)Image: colspan="2">Normal skin = 0Fontamelle: normal = 021Normal skin = 0abnormal = 1Image: colspan="2">Normal skin = 0specifyImage: colspan="2">Server active eczema = 3Kormal skin = 022Mild active eczema = 4SpecifyImage: colspan="2">Server active eczema = 5Normal: no infection = 023Image: colspan="2">Non-eczematous rash = 8Nose: no infection = 024Ante cubital fossa31Nose: no infection = 1Image: colspan="2">Non-eczematous rash = 8Nose: no infection = 1Image: colspan="2">SecifyNose: no infection = 1Image: colspan="2">SecifyNoter: normal = 026Multe33other = 2Trunk - back3933SpecifyImage: colspan="2">Tunk - frontAbdomen: normal = 12727abnormal = 12727abnormal = 127other absent = 028present = 129Other absent = 029abnormality: present = 129Treatment none = 0for eczema past = 146	557				Serial No.
Nose:no infection = 024 infection = 1Face (exz. perioral)30 Ante cubital fossaCervical glands:normal = 025 enlarged = 1Popliteal fossa31 wristChest:normal = 026 wheeze = 1 other = 2Ankle34 ankleSpecify abnormal = 126 other = 2Arm36 armAbdomen:normal = 027 abnormal = 127 Perioral37 and anormalRash:absent = 028 present = 128 PerioralNeckOther abnormality:29 present = 1Treatment none = 0 for eczema46	Length (cm) Head circumfe Fontanelle: specify Ears: Throat:	erence (cm) normal abnormal no infection infection no infection	= 0 = 1 = 0 = 1 = 0	21 21 22	Code as follows : Normal skin = 0 History of eczema = 1 Inactive eczema = 2 Mild active eczema = 3 Moderate active eczema = 4 Severe active eczema = 5 Non-eczematous rash = 8
derivitarnormal 0 0 glands:enlarged $= 1$ 0 enlarged $= 1$ 0 $Makle$ 34 Ghest:normal $= 0$ 26 $4nkle$ 35 $wheeze$ $= 1$ 0 $other$ $= 2$ $specify$ $abnormal$ $= 0$ 27 27 27 $abnormal$ $= 1$ $present$ $= 1$ $present$ $= 0$ 28 $present$ $= 0$ 29 $Treatment$ $none$ $none$ $= 0$ 46		no infection	= 0		Ante cubital fossa 31
wheeze= 1Iother= 2IspecifyImage: specify in the specified in	glands:	enlarged	= 1		Ankle 34 Hand 35
Abdomen:normal= 027 $beinp$ abnormal= 11Perioral41specifyPost aural43Rash:absent= 028present= 11Otherabsent= 029abnormality:present= 1 $present$ = 11 $present$ = 1 $present$ =	100	other	= 2		Leg 37 Trunk - front 38
Rash:absent= 028Neck44present= 1 \square Neck44Other abnormality:absent= 029Treatment none= 0present= 1 \square \square Treatment none= 0for eczema past= 1 \square 46	Ab domen:			27	Perioral 41
abnormality: present = 1 for eczema past = 1 46		absent		28	Neck 44
	abnormality:	present	= 1	29	for eczema past = 1 46

Appendix 3 b

FORM 6A

	Cold :	no = 0; yes = 1. nasal symptoms≥ 2 weeks = 2.	47
	Cough :	no = 0; yes = 1.	48
	Wheeze :	no = 0; yes = 1.	49
	Thrush :	no = 0; Yes = 1.	50
	Vomiting :	no = 0; yes < 2 weeks = 1; yes > 2 weeks = 2.	51
	Diarrhoea :	no = 0; yes < 2 weeks = 1; yes ≥ 2 weeks = 2.	52
	Nappy rash :	no = 0; yes = 1.	53
	Other rash :	no = 0, yes = 1.	54
		4 · · · · · · ·	1
Do you ever see	damp patches o	n the walls of any room?	1

During the last 3 months, has the baby had the following ?

no = 0; yes = 1. Do you ever see patches of mould on the walls of any room?

you ever see patches of mould on the walls of any room no = 0; yes = 1.

Appendix 4a

Baby's name	1-12 Serial No.
13-20 Weight (Kg x 100) Length (cm) Head circumference (cm) Fontanelle: normal = 0 21 abnormal = 1 Specify Ears: no infection = 0 22 infection = 1 Throat: no infection = 0 23	Details of rash Code as follows : Normal skin = 0 History of eczema = 1 Inactive eczema = 2 Mild active eczema = 3 Moderate active eczema = 4 Severe active eczema = 5 Non-eczematous rash = 8 (specify)
infection = 1	Face (exz. perioral)30Ante cubital fossa31Wrist32Pepliteal fossa33
glands: enlarged = 1	Ankle 34 Hand 35 Arm 36 Leg 37 Trunk - front 38
specify = 0 27 Abdomen: normal = 0 27 abnormal = 1 specify	Trunk - back39Scalp40Perioral41Post aural42
Kash: absemt = 0 28 present = 1	Axilla 43 Neck 44 Napkin area 4
Other absent = 0 29 abnormality: present = 1	Treatment none = 0 for eczema past = 1 46 present = 2

ALLERGY PREVENTION STUDY : EXAMINATION AT SIX MONTHS

FORM 6B

55

During the last 3 months, has the baby had the following ?

Cold	:	no = 0 yes = 1 nasal symptoms ≥ 2 weeks = 2.	47
Cough	:	no = 0, yes = 1.	43
Wheeze	:	no = 0; yes = 1.	49
Thrush	:	no = 0. yes = 1.	50
Vomiting		no = 0; yes < 2 weeks = 1; yes≥ 2 weeks = 2.	51
Diarrhoea	:	no = 0; yes≮2 weeks = 1; Yes≥ 2 weeks = 2.	52
Nappy rash	:	no = 0; yes = 1.	53
Other rash	:	no = 0; yes = 1.	54

During the last 6 months, has the baby been treated by a doctor? no = 0; yes = 1.

If so, what was the matter?

Results of skin tests . (code weal size; 8mm+ = 8; not done = 9)

56	57	58	59 6	60 61	62	63	64	65	66	67	68	69 70) 71	
ak	hinge	+	vi		Mile	1	1	Dry	1	1	T Egg		1	
	,list		her Lsl	kaulu j	D. pte	r.1 flow			nig: eout	-af	Tr	1	· Soy	a ·
	fee	111	5	mil	ĩ	f	ish		A	speg		Gra	us po lle	en

9

Appendix 5a

	ALLERGY PPEV	ENTION ST	CUDY :	EXAMINATION AT TWELVE MONTHS
Baby's name				Serial No.
Age			-	Card 5 Date :
Weight (Kg x length (cm) Head circumf Fontanelle:	erence (cm)	13-20	21	<u>Details of rash</u> Code as follows : Normal skin = 0 History of eczema = 1 Inactive eczema = 2
	abnormal	= 1		Mild active eczema = 3 Moderate active eczema = 4
specify				Severe active eczema = 5
Contract of Carlos	no infection infection		22	Non-eczematous rash = 8 (specify)
Throat:	no infection infection		23	
Nose:	no infection infection	= 0	24	Face (exc. perioral) 30 Ante cubital fossa 31
Cervical glanús:	normal enlarged		25 	Wrist 32 Popliteal fossa 33 Ankle 34
Chesti	normal wheeze other	= 0 = 1 = 2	26 	Hand 35 Arm 36 Leg 37
specify				Trunk - front 38
Abdomen:	normal abnormal	= 0 = 1	27	Trunk - back 39 Scalp 40 Perioral 41
specify				Post aural 42
Rash:	absent present	= 0 = 1	28	Axilla 43 Neck 44
Other abnormality specify	absent present	= 0 = 1	29 	Napkin area 45 Treatment none = 0 for eczema past = 1 46 present = 2
	2			

FORM 6C

Appendix 5b

During the last 6 months, has the baby had the following?

	Cold	:	no = 0;	yes = 1;				
			nasal s	ymptoms 3	2 weeks = 2.	*		47
							<u> </u>	
9	Cough	:	no = 0;	yes = 1.	а — а			43
				¥ 8				
	Wheeze	:	no = 0;	yes = 1.				49
	_		-	10000000001 10000			1 1	-
	Thrush	÷	no = 0;	yes = 1.				50
	Vomiting		$\mathbf{n}\mathbf{o} = 0$	yes < 2 w	mooks = 1;	.3		
	vour cring	•		weeks = 2.			Π	51
	10 C		100 -	Notico - L	53		·ł	51
	Diarrhoea	:	no = 0;	yes < 2 w	veeks = 1;		1010202	
			yes.≥ 2	weeks = 2.	no na			52
	Nappy rash	:	no = 0;	yes = 1.				53
							-	
	Other rash	:	No = $0;$	yes = 1.				54
			1					
8	treatment b				yes = 1.	• •		
	If so, what	wa	s the ma	tter?			<u> </u>	55
	<u> </u>							
Has the baby be	an admitted	to	hospital	at any tir				
has the baby be	en aumitieu		uospicui		yes = 1.			56
If go what	was the mat	or'	, .	,	,			50
Have you had an				ce you had	the baby?			
					yes = 1.	cat	Π	57
						dog		58
					ot	her mammal		59
				· . ·	birds	only other	\Box	60
					Ignore fist		•	
Does anyone liv	ing in the h	ous	e smoke	cigarettes?			· ·	
no = 0; me	other = 1; ϕ	oth	er perso	n, not moth	er = 2.			61
Have you had an	open coal f	ire	in the 1	house since	you had the	baby?	 1	÷
				no = 0	yes = 1.			62
			_	1 ×				
Number of perso	1200 CONTRACTOR (21) 10		131 Mar 141 162	9+ = 9:			H	53
Number of rooms	(incl. kitch	hen	only if	used for e	ating meals)		<u> </u>	64
Skin toota	45 26 67 69		70 71 7	10 70 74 75	76 77 76 70	80		
Skin tests :		1		1 1	76 77 78 79			
	the states of th	1.	I and the set	1 1 1	1 1 1	1		

Appendix 6 Questionnaire completed from 2nd-6th birthdays.

CONFIDENTIAL		
NameRef. No		
PLEASE PUT A RING ROUND THE CORRECT ANSWER LIKE THIS	YES	NO
 Have you consulted a doctor about him/her during the last 12 months? If so, what was the matter 	YES	NO
 Have you heard a wheeze coming from his/her chest during the last 12 months? (wheeze means a whistling sound whether high or low-pitched, and however fair 	YES	NO
3. Has he/she been treated for eczema by a doctor during the last 12 months?	YES	NO
f your address has changed, please give your new address:—		

This questionnaire was completed annually on about each childs 2nd - 6th bithdays.

Appendix 7- Age 7

ALLERGY PREVENTION STUDY	1
	1
Name	$\begin{array}{c} 6 \\ 7 \\ 1 \\ F = 2 \\ 7 \end{array}$
PLEASE PUT A RING AROUND THE RIGHT ANSWER LIKE THIS -	
Has your child ever suffered from any of the follow- ing? (Put a ring around the name of any disease(s) he/she has had).	9 N
BRONCHITIS ASTHMA HAY FEVER/ALLERGIC RHINITIS	8
ECZEMA PNEUMONIA/PLEURISY HEART TROUBLE	11
Have you consulted a doctor about him/her during the last 12 months?	
YES NO As. = 1, Other resp. = 2, Other = 3	14
If so, what was the matter?	
Have you heard a wheeze coming from his/her chest during the last 12 months? YES NO	15
(wheeze means a <u>whistling</u> sound, whether high or low-pitched, and however faint).	
If YES,	
Does it ever disturb his/her sleep? YES NO	16
Do any of the following bring on the wheeze?	
RUNNING WORRY EXCITEMENT COLDS	
ANIMALS e.g	17
FOODS/DRINKS e.g.	21
Any other factor	
Do colds usually go on to his/her chest? YES NO	23
Has he/she been treated for eczema by a doctor during	
the last 12 months? YES NO	24

13

Appendix 7b

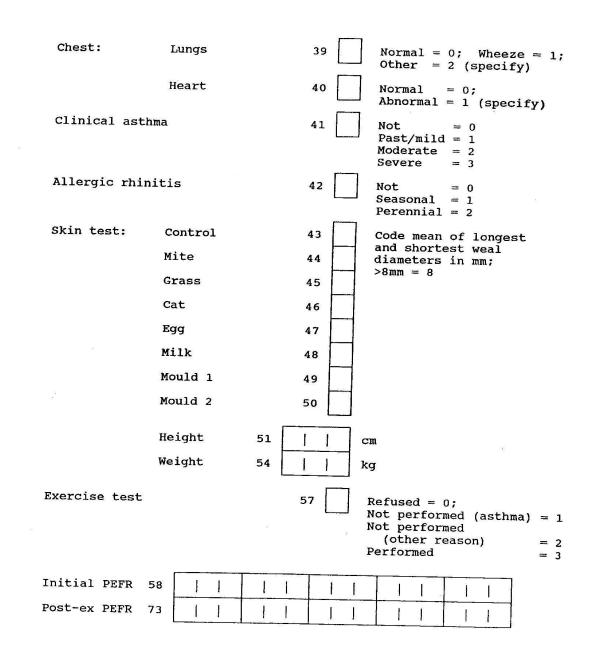
	2.		
Please record any treatment he/ the last 48 hours	she has taken during		
Bronchodilator Inh.=1, Ora	1=2, Both=3	25	
DSCG Steroid Inh.=1, Ora	1=2, Both=3	27	
What pets, if any, do you have? CAT DOG OTHER MAMMAL BIRD OTHER		28	
Do you ever see damp patches on room?	the walls of any YES NO	33	
Do you ever see patches of moul any room?	d on the walls of YES NO	34	
How many people living in the h	ouse smoke cigarettes?	35	
No. persons now living in house	? $9+ = 9: NK = 0$	36	
No. rooms (incl. kitchen if use	d for eating meals)? 9 + = 9 NK = 0	37	
Do you use gas for cooking?		38	
Is the house centrally heated? No = 0; Radiators = 1;	Ducts = 2	39	
Other forms of heating: Op	en coal fires	40	
cı	osed solid fuel stove	41	*
Ga	s fire	42	
El	ectricity	43	
ot	her	44	
Occupation of main wage earner		45	
		46 T]

Appendix 7c

		Seri	al no.		1					
		Card	5.					6		8
Name		Date	of bir	th 7			1	T	1	7
Address		mada	y's dat	e 13	H	-	$-\frac{1}{1}$	-†-	·	-
		Toua	y's uac	e 13						
Rash:	Face (exc. perioral	l) 19		Norma Histo			czei	na		= 0 = 1
	Antecubital fossa	20		(past Inact	yea	r)				= 2
	Wrist	21		Mild Moder	acti	ve	ecze		ema	= 3 = 4
	Popliteal fossa	22	Ц	Sever Rash,					1	= 5 = 6
	Ankle	23								
	Hand	24	Ц							
	Arm	25	Ц							
	Leg	26								
	Trunk-front	27	Ц							
	Trunk-back	28								
*	Scalp	29	Ц							
<u>a</u>	Perioral	30								
	Postaural	31						1		
	Axilla	32								
	Neck	33								
Inflammation:	Conjunct.	34		Norm=0;	Mil	d i	nj.=	=1;]	[nf]	.=2
	Throat	35		Norm=0;	Inf	1.=	1			
	Ears	36	· •	Norm=0;	SON	1=1;	Inf	1.=2	2	
	Nose/Cerv.gl.	37		Norm=0;	Inf	1./	Enla	argeo	1=1	

ALLERGY PREVENTION STUDY: EXAMINATION AT SEVEN YEARS

Appendix 7d



- 2 -

16

ALLERGY PREVENTION STUDY All answers will be treated as strictly confidential

Nar	me		Address		1
Dat	e of Bir	th			2
PLE	EASE P	UT A TICK BESIDE THE RIGHT ANSW	ERS LIKE THIS YES	Ø	
1	moi	re you heard a wheeze coming from your hths? heeze" means a <i>whistling</i> sound, whether high or i ES,		Yes 🗍 1 No 🗍 0	3
	a)	In the last 12 months, how often on a has your sleep been disturbed by whe	eezing? Less than one ni	Never 🗍 0 ght per week 🗍 1 hts per week 🗍 2	4
	b)	In the last 12 months, has wheezing e to limit your speech to one or two wor	ever been severe enough rds between breaths?	Yes 🗍 1 No 🗍 0	5
	c)	What do you think brings on the whee	ezing?		
		Smoke Perfumes D	Food 🗍	nimals 🗍 Drink 🗍	6 14
		(please specify)	
2	In the coug	e last 12 months, have you had a dry cou h associated with a cold or chest infectio	ugh at night, apart from a n?	Yes 🔲 1 No 🗍 0	15
3	in the staye	e last 12 months, how many days have yo d indoors because of chest trouble?	ou missed school or	None 🔲 0 1-3 🗍 1 4-7 🗍 2 8 or more 🗍 3	16
4	In the runny	last 12 months, have you had a problem or blocked nose, when you DID NOT ha	n with sneezing, or a ave a cold or the 'flu?	Yes 🗍 1 No 🗍 0	17
	If Yes				
	In the itchy-	last 12 months, has the nose problem be watery eyes?	een accompanied by	Yes 🗍 1 No 🗍 0	18
5	Have six mo	you <u>ever</u> had an itchy rash that was com onths?	ing and going for at least	Yes 🗍 1 No 🗍 0	19
	If Yes				
	a)	Have you had this itchy rash at any time		Yes 🗖 1 No 🗍 0	20
	b)	Has this itchy rash at any time affected			
		the folds of the elbows, behind the knew under the buttocks or around the neck,	es, in front of the ankles, ears or eyes?	Yes 🗍 1 No 🗍 0	21

Appendix 8b

6	Have you <u>ever</u> si	uffered from any of the follo	wing?				
	Asthma 🗖	Hay fever or Allergic Rhir	iitis 🗖	Eczema 🗖			22- 24
7	Have you <u>ever</u> be	een treated for eczema by a	doctor?	Yes 🗍 No 🗖	1 0		25
8	Have you <u>ever</u> sr	noked a cigarette?		Yes 🗖 No 🗖	1 0	2	26
	lf Yes, Do you smoke ciç	garettes every day?	á "	Yes 🗖 No 🗖	1 0	2	27
9	Does anyone else	living in your house smoke	cigarettes?	Yes 🗍 No 🗍	1 0	2	28
10		ths, have you taken any tre s, or inhalers) for chest trou			1 0	2	9
	If Yes, which of th	e following have you taken	and when?				
	Type of	treatment	(ti	When take			
			At fixed times every day	As needed when wheezy			led before ercise
Inhale	d Relievers: Ventolin, Bricanyl,	Salbutamol, Terbutaline	Π	П			
Inhaled	d Steroid Preventer Becotide, Flixotide Beclomethasone,		[°] O	ć (D)	·	·	 J
Inhaled	d Other Preventers Intal, Cromoglycat		٥			C	3
Give na	inhalers/Nebulisers ame(s)		٦	0	,e	C	5
				٥		C	J
	Tablets:		9	~		_	
	Prenisolone, Predr	nesol,		٥		E]
Other T Give na	, <i>,</i> ,		٥	٥		C]
			σ	σ			J

Appendix 9a- age 23

.

Date: _/_/ Time: _____

The Allergy Prevention Study-Patient Questionnaire Version 2 January 2006

Name	DOB
Address	E/O

Family History			
Mother	Hayfever 🗆	Eczema 🗆	Asthma 🗆
Father	Hayfever 🗆	Eczema 🗆	Asthma 🗆
Sibling Children	Hayfever 🗆	Eczema 🗆	Asthma 🗆
Children	Hayfever 🗆	Eczema 🗆	Asthma 🗆

Housing	Owner 🗆 Rented 🗆 Council 🗆
0	$Gas \sqcap Electric \sqcap None \sqcap$
Central heating?	
Cooking fuel	Gas 🗆 Electric 🗆 Oil 🗆 Other 🗆
Open fire?	Yes 🗆 No 🗆
Any damp?	Yes 🗆 No 🗆 DK 🗆
Do you have children?	Yes 🗆 No 🗆
House cohabitants?	Yes 🗆 No 🗇 Details
Pets?	Yes 🗆 No 🗆 Details
Employment history	Yes 🗆 No 🗆 Details

Appendix 9b

1.	Have you heard a wheeze (a whistling sound, whether high or low pitched, and however faint) coming from your chest in the last 12 months? Yes \Box No \Box
	If yes 1a) In the last 12 months how often has your sleep been disturbed by wheezing Never Less than one night a week One or more nights a week
	1b) In the last 12 months has wheezing ever been severe enough to limit your speech to one or two words between breaths? Yes□ No □
	1c) What do you think brings on the wheezing Cold □ Animals Smoke □ Perfume Perfume □ Food Other □ (please specify)
2.	In the last 12 months have you had a dry cough at night apart from a cough associated with a cold or chest infection? Yes No
	If yes 2a) Do you cough like this on most days for as much as 3 months each year? Yes □ No □
3.	Do you usually cough first thing in the morning in the winter? Yes No
	If Yes 3a) Do you cough like this on most days for as much as 3 months each year? Yes □ No □
4.	Do you usually cough during the day,or at night, in the winter? Yes No No
	If yes 4a) Do you cough like this on most days for as much as 3 months each year? Yes □ No □
5.	Do you usually bring up any phlegm from your chest first thing in the morning in winter? Yes No No

20

Appendix 9c

6.	Do you usually bring up any phlegm from your chest during the day, or night, in the winter? Yes No
	If Yes to Q5 or Q6 6a) Do you bring up phlegm like this on most days for as much as 3 months each year? Yes No
7.	In the past 3 years have you had a period of (increased) cough and phlegm for three weeks or more? Yes No
	If Yes 7a) Have you had more than one such period? Yes No
8.	Are you troubled by shortness of breath when hurrying on level ground or walking up a slight hill ? Yes No
9.	In the last 12 months how many days have you missed work or stayed indoors because of chest trouble? None 1-3 4-7 8 or more
10.	In the last 12 months have you had a problem with sneezing, or a runny or blocked nose, when you DID NOT have a cold or flu? Yes No
	If Yes. 10a) In the last 12 months has the nose problem been associated with itchy-watery eyes? Yes No
11.	Have you <u>ever</u> had an itchy rash that was coming and going for at least 6 months? Yes No
	If Yes 11 a) Have you had this itchy rash at any time in the last 12 months? Yes No
	 11b) Has this itchy rash at any time affected any of the following places: The folds of the elbows, behind the knees, in front of the ankles, under the buttocks or around the neck, ears or eyes? Yes No

12.	Have you <u>ever</u> suffered from any of the following Asthma Hay fever/allergic rhinitis Eczema
13.	Have you <u>ever</u> been treated for eczema by a doctor? Yes No
14.	Have you ever smoked cigarettes? Yes No
	If Yes 14a)Do you now smoke cigarettes every day? Yes No
	If Yes 14b)How much do you now smoke? 14c)How old were you when you started smoking?
	If No 14d)Have you ever smoked for as long as a year? Yes No
15.	Have you ever smoked any other form of tobacco eg cigars, pipe? Yes No If yes please give details?
16.	Does anyone else living in your house smoke? Yes No
17.	In the last 12 months have you taken any treatment for chest trouble? Yes No
	If Yes, please give following information
	Inhaled reliever Please list
	Inhaled steroid Please list
	Inhaled other preventers Please list
	Other inhalers/Nebulisers Please list
	Steroid tablets Please list
	Other tablets/medicine Give names

- Appendix 9e

18.	In the last 48 hours have you taken any treatment for chest trouble? Yes □ No □
	If Yes, which of the following have you taken and when?
	Inhaled reliever Please list
	Inhaled steroid Please list
	Inhaled other preventers Please list
	Other inhalers/Nebulisers Please list
	Steroid tablets Please list
	Other tablets/medicine Please list

Appendix 9f

Height	
--------	--

Weight

Skin Prick To	ests		
Neg control		Trees	
Pos control		Moulds	
Mites		Cat	
Grass		Dog	

Lung Function Tests				
Baseline				
FVC				
FEV1				
PF				

15mins post ventolin

Nitric Oxide level

8. Study instruments for 6/7 year olds

8.1 Instructions for completing questionnaire and demographic questions

Examples of instructions for completing questionnaire and demographic questions are given below. <u>The content of the questionnaires is fixed.</u> (see pages 72–73 for 'office use only' boxes example)

On this sheet are questions about your child's name, school, and birth dates. Please write your answers to these questions in the space provided.

All other questions require you to tick your answer in a box. If you make a mistake put a cross in the box and tick the correct answer. Tick only one option unless otherwise instructed.

Examples of how to :	mark questionnaires: Age 6 years
To answer Yes/No, put a appropriate box as per	
SCHOOL:	
TODAY'S DATE:	Day Month Year
CHILD'S NAME:	
CHILD'S AGE:	
CHILD'S DATE OF BIRTH:	years Day Month Year
(Tick all your answers	for the rest of the questionnaire)
Is your child a:	MALE FEMALE

Optional questions on ethnicity here

Appendix 10b

8.2	Core questionnaire for asthma				
8.2.1	Questionnaire for 6/7 year olds (strongly recommended)				
1	Has your child <u>ever</u> had wheezing or whistling in the chest at any time in the past?		Yes No		
	IF YOU HAVE ANSWERED "NO	D" PLEASE SKIP TO QUESTION	16		
2	Has your child had wheezing or whistling in the chest in the past 12 months? IF YOU HAVE ANSWERED "NO)" PLEASE SKIP TO QUESTION	Yes No		
3	How many attacks of wheezing has your child had in the past 12 months?	None 1 to 3 4 to 12 More than			
4	In the past 12 months, how often, on average, has your child's sleep been disturbed due to wheezing?				
		Never woken with wheezing Less than one night per week One or more nights per week			
5	In the past 12 months, has wheezin ever been severe enough to limit yo child's speech to only one or two words at a time between breaths?		Yes No		
6	Has your child <u>ever</u> had asthma?		Yes No		
7	In the past 12 months, has your child's chest sounded wheezy during or after exercise?		Yes No		
8	In the past 12 months, has your child had a dry cough at night, apart from a cough associated with a cold or chest infection?	i.	Yes No		

7. Study instruments for 13/14 year olds

7.1 Instructions for completing questionnaire and demographic questions

Examples of instructions for completing questionnaires and demographic questions are given below. <u>The questionnaire content is fixed</u>. (see pages 72-73 for 'office use only' boxes example)

On this sheet are questions about your name, school, and birth dates. Please write your answers to these questions in the space provided.

All other questions require you to tick your answer in a box. If you make a mistake put a cross in the box and tick the correct answer. Tick only one option unless otherwise instructed.

Examples of how to	mark questionnaires: Age 13 years
To answer Yes/No, put a appropriate box as per	
SCHOOL:	
TODAY'S DATE:	Day Month Year
YOUR NAME:	
YOUR AGE:	
YOUR DATE OF BIRTH:	years Day Month Year
(Tick all your answers	for the rest of the questionnaire)
Are you:	MALE FEMALE
Optional questions on e	thnicity here

26

Appendix 10d

7.2	Core questionnaire for asthma			
7.2.1	Questionnaire for 13/14 year olds			
1	Have you <u>ever</u> had wheezing or whistling in the chest at any time in the past?		Yes No	\square
	IF YOU HAVE ANSWERED "NO	" PLEASE SKIP TO QUESTION	16	
2	Have you had wheezing or whistling in the chest in the past 12 months? IF YOU HAVE ANSWERED "NO	" PLEASE SKIP TO QUESTION	Yes No	\square
3	How many attacks of wheezing have you had	None 1 to 3		B
	in the past 12 months?	4 to 12 More than	12	
4	In the past 12 months, how often, or your sleep been disturbed due to wh			
5	In the past 12 months, has wheezing ever been severe enough to limit you speech to only one or two words at a time between breaths?		Yes No	\square
6	Have you <u>ever</u> had asthma?		Yes No	
7	In the past 12 months, has your chest sounded wheezy during or after exercise?		Yes No	\square
8	In the past 12 months, have you had a dry cough at night, apart from a cough associated with a cold or chest infection?		Yes No	\square

Module 3.2: Skin prick tests for atopy

Aims

- To provide an objective measure of atopy for comparisons within and between centres. "Atopy" may be defined as skin test reactivity to one or more of the following allergens: house dust mites (*Dermatophagoides pteronyssinus* or *Dermatophagoides farinae*); cat fur; mixed grass pollen; mixed tree pollen; and the outdoor mould genus *Alternaria*.
- To compare the prevalence and degree of sensitivity to individual allergens which are ubiquitous within and between centres (e.g. mites, cat, Alternaria).

Methods

When selecting an appropriate skin prick test method for ISAAC several criteria have been applied: reproducibility under field conditions; simplicity of application; safety; acceptability; quality control; suitability for all ISAAC age groups; low cost; and worldwide availability.

The ALK lancet has been chosen for a number of reasons. Reproducibility and precision with both histamine and allergen extracts has been shown to be good [1]. The application is simple, safe and accepted by children, parents and field workers. There is a good body of knowledge in the medical literature on this skin prick test method and it has already been applied in many surveys. The ALK lancet is available throughout the world.

The "core" allergen extracts to be tested on the *left* forearm are:

Histamine 10 mg/ml (positive control)	Diluent (negative control)
D. pteronyssinus	D. farinae
Cat	Alternaria tenuis
Mixed grasses	Mixed trees

In addition, each centre may add up to eight allergens of their choice by testing them on the *right* forearm. The local relevance of cockroach, artemesia, olive and ragweed should be considered. All of these extracts should be purchased from ALK Denmark. In addition, local allergens that are most prevalent in the respective study area should be included and purchased if possible from ALK, or if not available from ALK, from other companies.

All extracts and the control solutions should be obtained from ALK Laboratories (full address listed on page 65). The allergen extracts are highly standardised and can be delivered throughout the world, including the USA. Histamine 10 mg/ml has been chosen as a positive control solution

because of better reproducibility and precision than alternative positive control solutions [2]. The grass extract is a mixture of commonly occurring grasses in central Europe, i.e.: *Dactylis glomerata*, *Lolium perennae*, *Festuca pratensis*, *Poa pratensis*, *Phleum pratense* and *Avena eliator*. The tree extract is a mixture of commonly occurring tree pollen in central Europe, i.e.: *Betula verrucosa* (birch), *Alnus glutinosa* (alder) and *Corylus avellana* (hazel).

There is a circadian rhythm in the size of skin prick reactions to allergens and histamine [3], so all skin prick tests should be performed in the morning hours (08:00 to 13:00, local time). The site of testing should be free of eczema. An ALK tape with numbers indicating the sequence of allergen extracts is placed in the middle of the volar aspect of the left forearm, 3 cm distal to the elbow crease. One drop of each skin prick solution is placed on the forearm in the above order, on the left and right sides of the tape, respectively. A separate ALK lancet is pricked vertically through each drop with firm pressure. All drops and the tape are removed immediately after the pricks taking care not to contaminate prick points with a different extract.

Reactions to each skin test solution are measured 15 minutes after the pricks. The contours of each wheal are outlined with a fine filter tip pen. The contours are then transferred to the record sheet by means of translucent tape. The size of each wheal is documented as the mean of the longest diameter (a) and the diameter perpendicular to it at its mid-point (b): i.e. (a+b)/2. Measurements of each diameter are made to the nearest millimetre above.

In dark skin wheals can be recognised more easily under strong oblique light and also by palpating the skin. In persons who spend much time outdoors the thickening of the skin may limit the ability to detect skin prick reactions.

Training

Field workers should be trained before starting the survey, and their precision retested in the middle and at the end of the survey, since the technique of individual fieldworkers may change over time. Reproducibility should be tested as follows at the start of the survey. At least three series of 16 skin prick tests with histamine 10 mg/ml should be performed by each field worker on the volar surface of the forearm of a volunteer until the coefficient of variation (standard deviation as a percent of the mean) of the last series is less than 20%. Half way through the survey and at the end of fieldwork, each field worker should perform two further

series of 16 skin prick tests with histamine 10 mg/ml on the volar forearm of a volunteer. All results should be documented separately for each fieldworker on the "training" record sheets.

Validation

Because of difficulties in standardizing the performance of different field workers, validation studies using serum IgE measurements (ISAAC module 3.5) are highly recommended in a subsample of children. Where possible, multi-centre comparisons should adopt a cross-over allocation of fieldworkers to the different study areas, so that approximately equal numbers of children are tested by each observer in each centre. Otherwise, it may become impossible to disentangle differences in the performance of different fieldworkers from real differences in the prevalence of skin test reactivity in the comparison areas.

Safety

Slight physical discomfort may result from the prick and itchiness of the larger wheals. Systemic allergic reactions have not been reported with prick testing despite extensive use in epidemiological surveys. Among over 16,000 adults and children tested in the United States NHANES II survey, six subjects fainted after prick testing, compared to 26 faints after venipuncture [4]. Reviews of deaths occurring from immunotherapy and skin testing in the USA found no fatalities that could be attributed to prick, puncture or scratch testing in the absence of intradermal tests or desensitisation immunotherapy [5, 6]. Systemic allergic reactions occur rarely (0.02%) with *intradermal* skin testing among allergic patients [7] but this technique will not be used in ISAAC.

References

- Nelson HS, Rosloniec DM, McCall LI, Ilké D. Comparative performance of five commercial prick skin test devices. J Allergy Clin Immunol 1993; 92: 750-756.
- Illi S, Garcia-Marcos L, Hernando V, Guillen JJ, Liese A, von Mutius E. Reproducibility of skin prick test results in epidemiological studies: a comparison of two devices. *Allergy* 1998; 53: 353-358.
- Taudorf E, Malling HJ, Laursen LC, Lanner A, Weeke B. Reproducibility of histamine skin prick tests. Inter- and intra-observer variation using histamine dihydrochloride 1,5 and 10 mg/ml. *Allergy* 1985; 40: 344-349.

- Dreborg S (ed). Skin tests used in type I allergy testing. Position paper prepared by the sub-committee on skin tests of the European Academy of Allergology and Clinical Immunology. *Allergy* 1989; 44[Suppl]: 22-30, 52-59.
- Turkeltaub PC, Gergen PJ. The risk of adverse reactions from percutaneous prick-puncture allergen skin testing, venipuncture and body measurements: data from the second National Health and Nutrition Examination Survey 1976-1980 (NHANES II). J Allergy Clin Immunol 1989; 84: 886-890.
- Lockey RF, Benedict LM, Turkeltaub PC, Bukantz SC. Fatalities from immunotherapy and skin testing. J Allergy Clin Immunol 1987; 79: 660-677.
- Reid MJ, Lockey RF, Turkeltaub PC, Platts-Mills TA. Survey of fatalities from skin testing and immunotherapy. J Allergy Clin Immunol 1993; 92: 6-15.
- Lin MS, Tanner E, Lynn J, Friday GA (Jr.). Nonfatal systemic allergic reactions induced by skin testing and immunotherapy. *Ann Allergy* 1993; 71: 557-562.

Contact address

PD Dr med Erika von Mutius Kinderklinik der Universität im Dr von Hauner'schen Kinderspital Lindwurmstraße 4 D-80337 München 2 GERMANY

Tel:	(49) 89 5160 2709
Fax:	(49) 89 5160 4452
E-mail:	erika.von.mutius@kk-i.med.uni-muenchen.de

Equipment checklist

- ALK allergen extracts
- Tray for allergen bottles
- ALK skin prick lancets
- ALK tape
- Swabs or tissues
- Sharps disposal container
- Felt tip pen (e.g. Edding 1800 profipen 0.5)
- Alarm clock
- Ruler
- Record sheets

FIELD MANUAL FOR SKIN PRICK TESTING

Allergen solutions

Inner side:

Outer side:

- 1. Positive control (histamine)
- 3. D. Pteronyssinus
- 5. Cat
- Mixed grasses

2. Negative control (glycerin)

- 4. D. farinae
- 6. Alternaria tenuis
- Mixed trees
- Perform these eight tests on the left forearm. Use a similar technique on the right forearm to test other allergens of local interest.
- Place the allergens on the tray in the same order as they are put on the forearm.
- Store allergen solutions in a refrigerator between test sessions.

Applying the solutions

- Check that the skin of the forearm is free of eczema. The test should not be performed on inflamed or broken skin.
- Place the left arm palm upwards on the table in front of the examiner.
- Paste a prenumbered ALK tape onto the left forearm, in the middle and with the "++" mark 2 cm from the elbow.
- Open the packaging of the ALK lancets before doing the test. They should be placed ready to be taken out of the package with one hand.
- Open the bottles with the allergen solutions.
- Put one drop of each allergen on the left or right side of the tape. Do this
 always in the same sequence. Do not use too much allergen and take
 care that the different allergens do not run together or run off the arm.
- Put the bottle back to its position on the tray. Do not change the order of the bottles.
- Always start applying allergens on the inner side, working from top (elbow) to bottom. The numbers 1, 3, 5, 7 on the tape mark the distance (1 cm) between the allergens. Apply allergens numbered 2, 4, 6, 8 on the outer side also from top to bottom.
- The drops of solution 1 and 2, 3 and 4, 5 and 6, as well as 7 and 8 are now next to each other, at the same height on the left and right side of the tape, respectively.

Performing the prick test

- Always use a new ALK lancet for each allergen.
- Prick the ALK lancet for 2 seconds vertically through the drop into the skin using firm pressure.
- Put the used lancets into the disposable container.
- After pricking wipe the allergens off without mixing them. Use a clean swab or tissue and wipe away from the tape towards the outside of the arm.
- Set the alarm clock for 15 minutes.
- · Close the allergen bottles with their own coloured caps.

Reading the reaction

- After 15 minutes outline the contours of the wheal with a thin felt-tip pen (e.g. Edding 1800 profipen 0.5). Do not spread the skin. Hold the pen vertically. Ensure adequate lighting.
- The contour should be drawn at the outside of the wheal. If there is no reaction mark that non-reactive position with a little dot.
- Write "I" on the skin at the top of the inner side and "O" at the top of the outer side, near the "++" mark on the tape.
- Remove the prenumbered tape.
- · Paste a transparent tape onto the wheals to transfer the contours.
- Press the tape onto the skin to make sure that the whole contour is transferred to the sticky side.
- · Remove the tape from the skin and paste it into the record sheet.

Measurement of each wheal

- Record measurements in millimetres, rounded to the next higher integer, using a flexible plastic ruler (e.g. Mérieux multitest).
- Always measure the inside of the felt-tip pen contour.
- Identify and measure the longest diameter first.
- Then drop a perpendicular line through the middle of the longest diameter and measure the length of this line.
- Calculate the mean of the two diameters.

SKIN PRICK TEST RECORD SHEET

ID number: _____ Area number: _____

Date: ____/___/____

Fix tape A here Fix tape B here

Field worker number: _____

VOLAR LOWER LEFT ARM

Tape A Tape B

1. +ve control 2. -ve control

3. D. Pteronyssinus 4. D. farinae

5. Cat 6. Alternaria tenuis

7. Mixed grasses 8. Mixed trees

DIAMETERS MEASURED TO THE NEAREST WHOLE MILLIMETRE:

1. Positive control		2. Negative control		
Max diam (a)	Min diam (b)	Max diam (a)	Min diam (b)	
	nyssinus	4. D. fi	rinae	
Max diam (a)	Min diam (b)	Max diam (a)	Min diam (b)	
5. C	at	6. Alterna	ria tenuis	
Max diam (a)	Min diam (b)	Max diam (a)	Min diam (b)	
7. Mixed grasses		8. Mixed trees		
Max diam (a)	Min diam (b)	Max diam (a)	Min diam (b)	