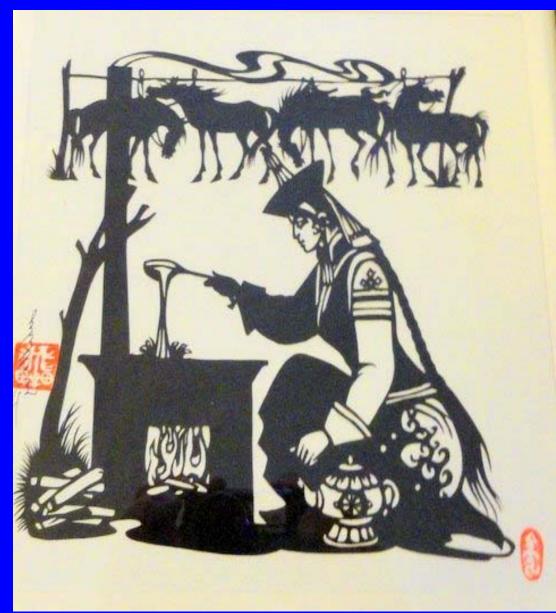
The Unfinished Health and Research Agendas of Incomplete Combustion Particles

Kirk R. Smith Tyler Laureate 2012 Professor of Global Environmental Health University of California, Berkeley

Environmental Mutagens in Human Populations, International Conference Doha, Qatar March 29, 2012



300-400 thousand years ago, hearths became a regular feature in human habitation

The skill that many anthropologists use for defining the point when humanity switched from pre-human to human conditions was learning to control fire (e.g., Levi-Strauss 1969).

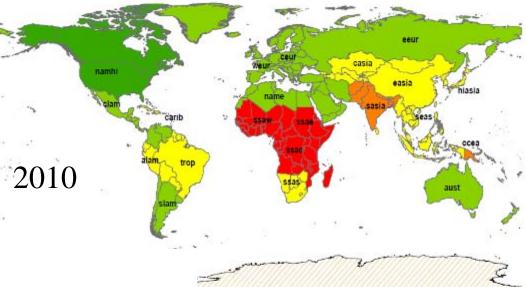
By this definition, cooking is the oldest task in human history.

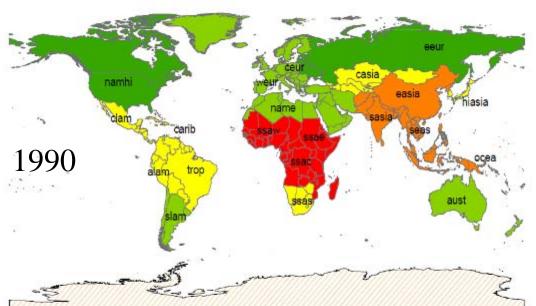
How long ago was this?

"On the earliest evidence for habitual use of fire" Roebroeks and Villa, PNAS, 2011

The three major solid fuels

Households using biomass or coal to cook



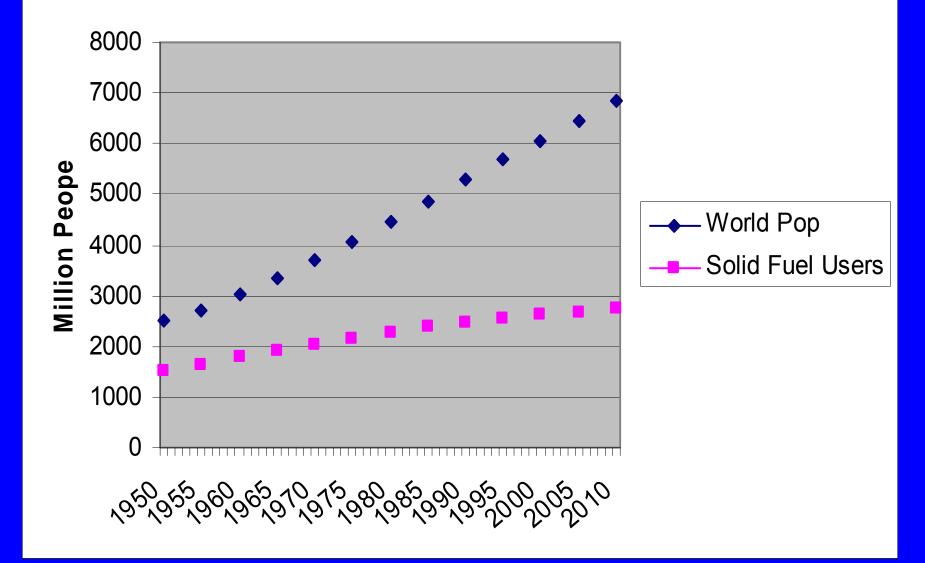


% of HH Exposed to HAP



Comparative Risk Assessment (CRA) 2012

World Population Using Solid Fuels



Woodsmoke is natural – how can it hurt you?

Or, since wood is mainly just carbon, hydrogen, and oxygen, doesn't it just change to CO_2 and H_2O when it is combined with oxygen (burned)?

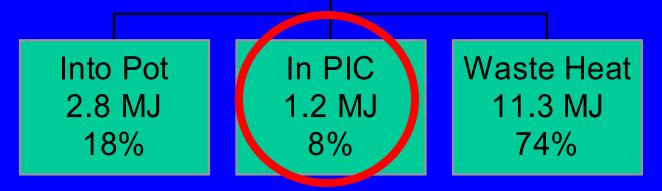


Reason: the combustion efficiency is far less than 100%

Energy flows in a well-operating traditional wood-fired Indian cooking stove

A Toxic Waste Factory!!

Typical biomass cookstoves convert 6-20% of the fuel carbon to toxic substances



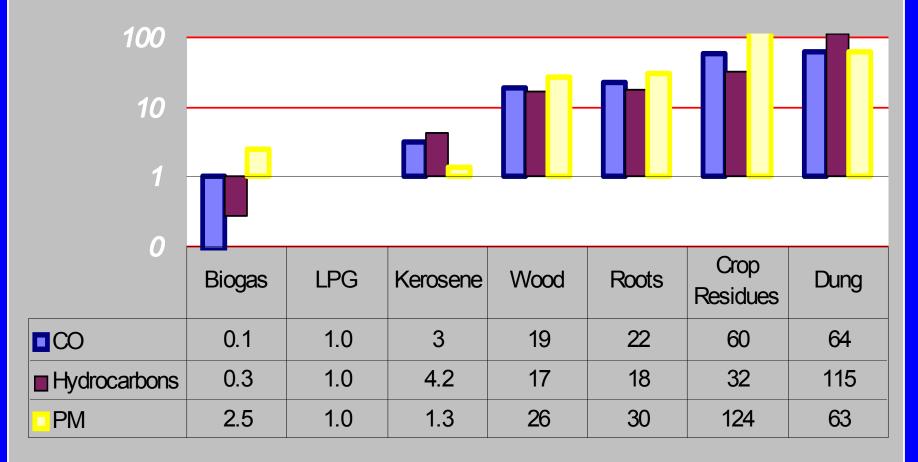
PIC = products of incomplete combustion = CO, HC, C, etc.

Source: Smith, et al., 2000

Toxic Pollutants in Biomass Fuel Smoke from Simple (poor) Combustion

- Small particles, CO, NO₂
- Hydrocarbons
 - 25+ saturated hydrocarbons such as *n*-hexane
 - 40+ unsaturated hydrocarbons such as 1,3 butadiene
 - 28+ mono-aromatics such as benzene & styrene
 - 20+ polycyclic aromatics such as $benzo(\alpha)pyrene$
- Oxygenated organics
 - 20+ aldehydes including *formaldehyde* & *acrolein*
 - 25+ alcohols and acids such as *methanol*
 - 33+ phenols such as *catechol* & *cresol*
 - Many quinones such as *hydroquinone*
 - Semi-quinone-type and other radicals
- Source: Naeher et al, *J Inhal Tox*, 2007
- Chlorinated organics such as *methylene chloride* and *dioxin*

Health-Damaging Pollutants per Unit Energy Delivered Ratio of Emissions to LPG



CO Hydrocarbons PM

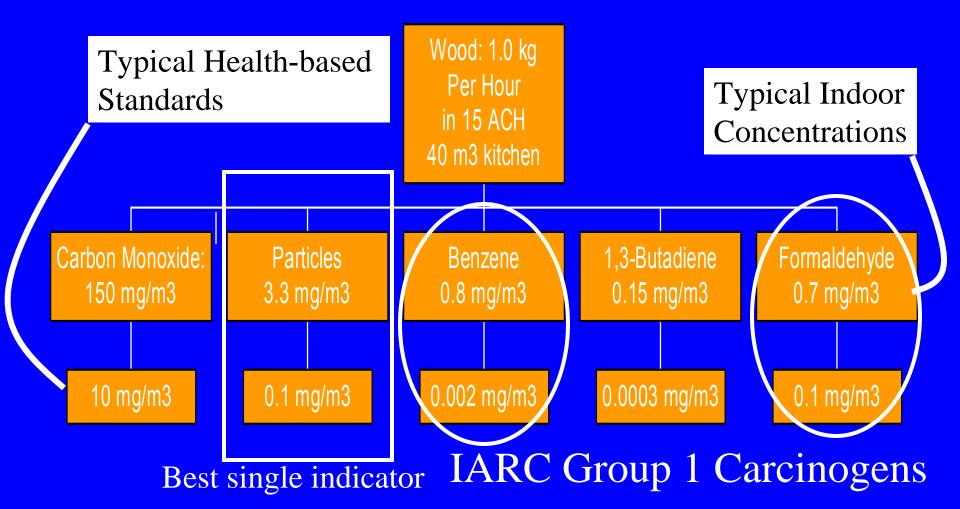
Smith, et al., 2005

First person in human history to have her exposure measured doing the oldest task in human history

Emissions, yes, but what about exposures?

> Kheda District, Gujarat, 1981

Health-Damaging Air Pollutants From Typical Woodfired Cookstove in India.



WORLD HEALTH ORGANIZATION INTERNATIONAL AGENCY FOR RESEARCH ON CANCER



IARC Monographs on the Evaluation of Carcinogenic Risks to Humans

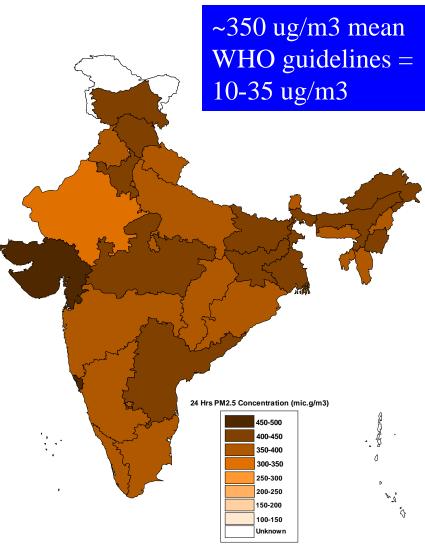
VOLUME 95

Household Use of Solid Fuels and High-temperature Frying

Published in 2010, but conducted in 2006. Woodsmoke found to be 2A – probable human carcinogen (Not Group #1 due to weak epidemiological evidence) Exposure Model for India based on measurements in ~1000 households

Estimated PM2.5 for <u>only</u> solid-fuel-using households

Household Air Pollution Comparative Risk Assessment, 2012



ALRI/ Pneumonia

Diseases for which we have epidemiological studies

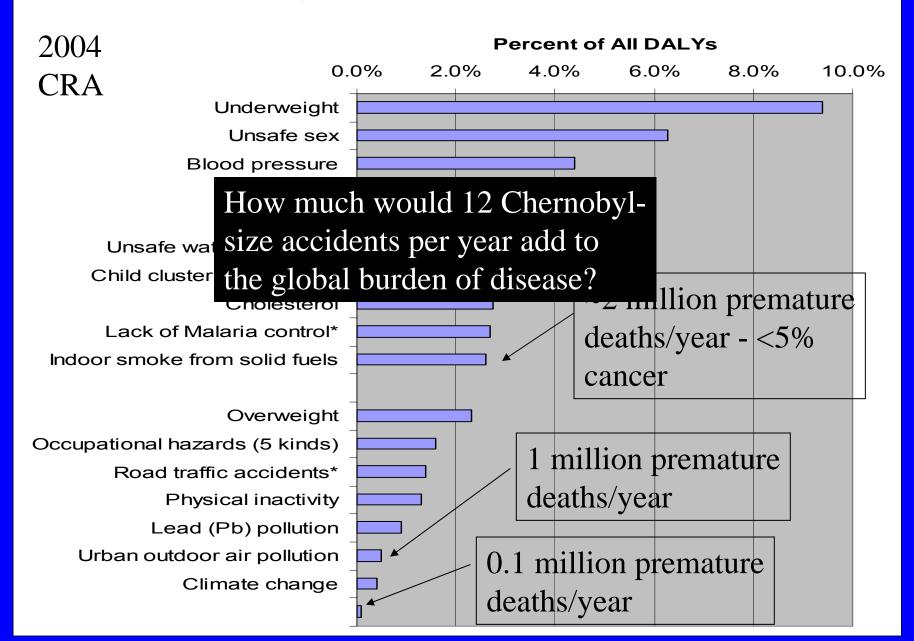
COPD Lung cancer (coal)



These three diseases were included in the 2004 Comparative Risk Assessment Managed and published by WHO

First ever comprehensive risk assessment with consistent rules of evidence and common databases

Global Burden of Disease from Top 10 Risk Factors plus selected other risk factors



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which makes which have allow pairs	n (thousands)	6224984.81		622741.70	817442.87	6633668.38	42707129	166160.20	30443.11	26143.24	3131052.05	301091.79	5892229.34	782026.21	646519.90	427846.34	181906.42	Ite
	002: DALYs by age, sex and cau	se for the	year 200	Z														
WORL	D		-				10-1-									F		
Code	Cause	Total	0-4	5-14	15-29	30-44	Male 45-59	60-69	70-79	80+	Total	0-4	5-14	15-29	30-44	Female 45-59	60-69	
U000	All Causes	1490125643		56142418	141637847	130501742	115374678	60641455		and there are the	772912264	C. Constant reconstant	54559890	141633214		90621463	53460940	40
	I. Communicable, maternal, perinatal	1430123643	222552013	30142410	141631041	130301/42	115314610	60641433	36223210	9832834	112312204	211279454	54553030	141033214	100251333	30621463	53460340	40
10.000	and nutritional conditions	610319230	*****	20979812	29721889	36148775	16631072	5676094	2913759	849889		******	25248712	59740290	35979204	11499502	4476964	25
U002	A. Infectious and parasitic diseases	350332571	87977510	12810197	25516201	33372707	14085712	3710016	1519252	315539	179307135	89749119	13534884	31549467	23070592	8616676	2658113	14
U003	1. Tuberculosis	34735908	839261	782431	5497848	7335503	4868489	1832663	658401	90612	21905208	738745	821735	4163278	3822011	2041939	833205	\$
U004 U005	2. STDs excluding HIV	11347067	1521528 1039452	26245 2292	1080828 151945	660600 285546	423721 354596	103004 99162	31641 30394	7415 6987	3854983 1970375	1726606 1265722	97221 3432	4158088	1147558 262487	260537 107605	71612 42492	
U005	a. Syphilis b. Chlamydia	4200039 3571404	1033452	7209	241593	50929	304036	136	30334	0307	302234	1260722	69338	521758 2635460		107605	42432	
U007	c. Gonorrhoea	2265159	462945	10559	£02964	201215	666A	719	01	2	1472160	447709	24222	909222	422675	6262	1616	
U008	d. Other STDs						- с г	1			- 4 - 1-				22947	37012	13420	
U009 U010	3. HIV/AIDS 4. Diarrhoeal diseases		GIO	Dal	Bura	den	OT L	JISE	ase	e Di	atab	ase			13746555 545936	2624463 373258	230670 201900	
U011	5. Childhood-cluster dise														53966	19174	4837	
U012	a. Pertussis		and		mn	arat	ivo	Die		200	ssm	ont			0	13	0	
U013	b. Poliomyelitis		апо		πр	aral	гие_			53 E	2211	EIII_			16931	3822	615	
U014 U015	c. Diphtheria						10	<u> </u>							386	572	29 0	
U015 U016	d. Measles e. Tetanus			-VV c	orld	Hea	alth_	Orc	an	zat	lon_				29 36620	30 14737	0 4192	
U017	6. Meningitis*								J M M						167759	110215	48775	
U018	7. Hepatitis B														115391	110437	36630	
U019	Hepatitis C														53705	61307	30322	
U020	8. Malaria														453207	280704	99111	
U021 U022	 Tropical-cluster disea: a. Trypanosomiasis 			Re	ing	con	nnle	toly	/ IIr	dat	bet				593336 79329	496341 46509	75537 3694	
U022	b. Chagas disease				in ig	COL	pic	JUIS	/ up	ua	LUU				51644	48058	15987	
U024	c. Schistosomiasis					0	040								128589	58850	17056	
U025	d. Leishmaniasis					or 2	UIZ	rei	eas	e					67203	30097	11647	
U026	e. lymphatic filariasis														206680	266630	15772	
U027 U028	f. Onchocerciasis 10. Leprosy														59892 15828	46198 9135	11380 4389	
U029	11. Dengue														10408	6396	2644	
U030	12. Japanese encephalitis									11 41					26926	7282	2513	
U031	13. Trachoma			FO	ho	use	noic	l alí	- DO	IIUI	lon:				517423	559520	344796	1
U032	14. Intestinal nematode in														5192	5921	3469	
U033 U034	a. Ascariasis b. Trichuriasis	ew oi	utoo	mo	octi	mol		220	od /	on	moto			00	161 388	21 433	111 238	
U035	c. Hookworm disease		uluu		C 31	IIIa	.621	Jas	eu '			a-an	arys	50	4212		230	
U036															432		397	
U037	Other infectious disease	LRI,	TCO	PD		na (Jan	cer	(tro	m	olom	ass	als	$\overline{\mathbf{D}}$	1795391	1650048	667703	
U038				- 2,		-9		-					are		1481589	1357246	1460806	13
U039 U040	1. Lower respiratory infect 2. Upper respiratory infec			Ce	tore	nota	C	rdi							1434738 45228	1329549 26339	1425397 34405	
U040	3. Otitis media				atara		$, \mathbf{b}_{\mathbf{c}}$	IT UI	Jva	<u>500</u>					40228	26333	1004	
U042	C. Maternal conditions	00001000									•		LLOUJL	20000100	9947305	403615	19	
U043	1. Maternal haemorrhage	4437585	0	0	0	0	0	0	0	0	0	50	· · · · · · · · · · · · · · · · · · ·	2359228		133887	0	
U044	2. Maternal sepsis	6903085	0	0	0	0	0	0	0	0	0	0	144	5204511	1625431	72999	0	
U045 U046	3. Hypertensive disorders* 4. Obstructed labour	2162701	0 0	0 0	0 0	0	0	0	0	0	0	0	975 0	1398598	722909	40216 12162	0 0	
U046 U047	4. Obstructed labour 5. Abortion	3048291 4652171	U N	U 0	U N	U N	U N	U N	U	U 0	U N	U 0	U 223411	2241561 3721304	794568 705860	12162	U 0	
U048	Other maternal conditions	12427759	0	0	0	0	0	0	0	0	0	0	1970	8128555		142755	0 19	
U049	D. Perinatal conditions*	97335086	53209265	1343	1031	347	89	12	9	0		44121066	1195	498		34	29	
U050	1. Low birth weight	46334234	25061999	52	13	20	0	9	0	0	25062092	21272111		0	0	0	0	
U051	2. Birth asphyxia and birth trauma	34445758		790	302	73	33	3	0	0	19354204	15090851	573	107		15	0	
U052 U053	Other perinatal conditions E Nutritional deficiencies	16555094 34416632	8794262 10258276	502 1921013	716 1793247	253 1025783	56 698252	0 230984	9 132831	0 39983	8795799 16100369	7758104 10385030	591 2582483	392 2149411		19 1121930	29 357998	
U053 U054	1. Protein-energy malnutrition	344 IBB 32 16910328	7556012	560106	156056	69278	103279	230984 71114	45451	39983 18671	8579966	7350453		66437		81418	397998 70683	
U055	2. Iodine deficiency	3519322	1283895	471857	528	748	1215	641	236	50		1305105	450178	1681		1287	439	
U056	 Vitamin A deficiency 	792562	257306	84825	3330	6358	7864	3519	983	100		320394	84034	10506			Full Scree	-
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ALRI/ Pneumonia

Low birth weight

Stillbirth

Diseases for which we now have epidemiological studies

COPD

Lung cancer (coal)

Lung cancer (biomass)

Blindness (cataracts, opacity)

CV disease Blood pressure ST-segment

These additional diseases will be included in the 2010 Comparative Risk Assessment – being published 2012

In addition, using evidence from other exposure sources, CVD will be included

There is epi evidence for these other diseases, but considered insufficient to include in the 2010 Comparative Risk Assessment



Burns and the health/safety impacts of fuel gathering

Tuberculosis ALRI

Other cancers (cervical, NP, upper airway)

Asthma?

Cognitive

Impairment

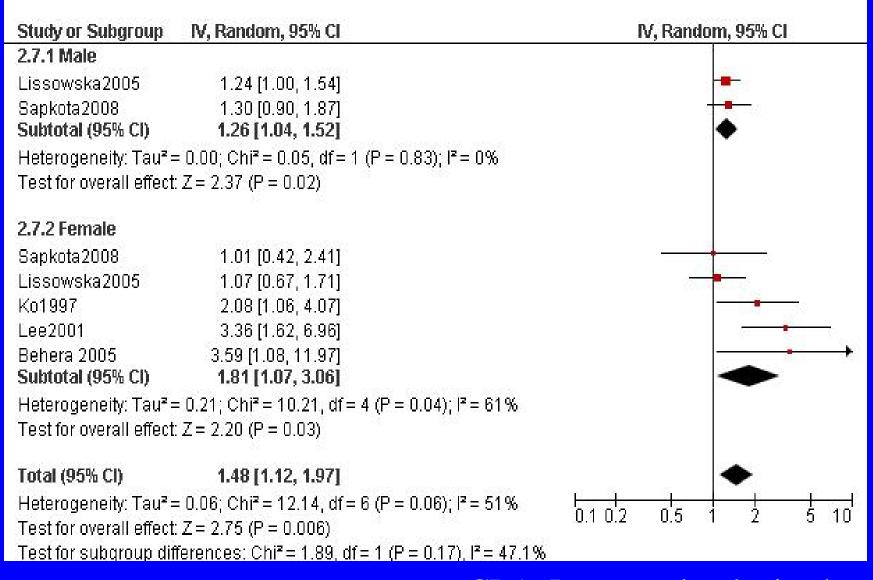
Birth defects

Systematic Review and Meta-analysis of lung cancer and household

coal combustion

study	studyyear o	casenum	contnum		ES (95% CI)
Africa Sasco(2002) Subtotal (I-square	1996-1998 1 d = .%, p = .)	118	235		0.74 (0.17, 3.14) 0.74 (0.17, 3.14)
Europe Lissowska(2005) Subtotal (I-square	1998-2001 2 d = .%, p = .)	2861	3118	\$	1.13 (0.93, 1.38) 1.13 (0.93, 1.38)
North America Wu(1985) Subtotal (I-square	1981-1982 2 d = .%, p = .)	220	220		2.30 (0.96, 5.50) 2.30 (0.96, 5.50)
India Gupta(2001) Sapkota(2008) Subtotal (I-square	1995-1997 2 2001-2004 7 d = 4.4%, p = 0	793	525 – 718		1.52 (0.33, 6.98) 3.76 (1.64, 8.63) 3.02 (1.42, 6.46)
Mainland China an Wu-Williams(1990 Sun(1991) Chengyu(1992) Ger(1993) Lin(1996) Dai(1996) Luo(1996) Ko(1997) Hao(1998) Huang(1999) Wu(1999) Lan(2000) Lee(2001) Kleinerman(2002) Sun(2002) Lu(2003) Liang(2004) Galeone(2008) Lan(2008) Subtotal (I-square) 1985-1987 1985-1987 1985-1987 1990-1991 1990-1991 1986-1992 1985-1990 1992-1993 1990-1991 1992-1993 1993-1996 1993-1996 1993-1998 1994-1998 1996-1999 2001-2002 1987-1990 2001-2002 1985-1990 4	418 135 131 161 122 120 102 117 220 122 258 122 258 122 258 122 258 122 258 122 258 122 228 122 228 122 228 122 228 122 228 122 228 132 228 132 228 132 228 132 228 132 228 132 132 132 132 132 132 132 132 132 132	959 398 135 524 161 122 120 306 117 440 244 258 122 805 1724 618 445 152 436 498		 1.30 (0.99, 1.70) 2.26 (1.53, 3.33) 1.59 (1.22, 2.07) 1.44 (0.44, 4.69) 2.08 (0.85, 5.08) 3.24 (1.05, 9.94) 4.70 (1.29, 17.18) 6.00 (5.07, 7.10) 1.30 (0.29, 5.80) 1.99 (1.16, 3.43) 1.92 (1.40, 2.62) 1.58 (0.89, 2.80) 2.40 (1.31, 4.40) 2.10 (1.19, 3.70) 1.29 (1.03, 1.61) 2.20 (1.25, 3.86) 3.44 (1.38, 8.57) 2.02 (1.20, 3.39) 2.19 (1.08, 4.46) 7.40 (4.18, 13.10) 2.28 (1.66, 3.13)
Overall (I-squared	= 90.4%, p =	0.000)			2.16 (1.62, 2.90)
			Odds ra	atio	

Lung Cancer: Biomass vs. clean fuel



CRA, Imran et al. submitted

Other outcomes

- Intriguing but with insufficient evidence to date to include as primary outcomes. Among them are:
 - Cognitive impacts in children
 - Cervical cancer



Contents lists available at SciVerse ScienceDirect

NeuroToxicology



Neurodevelopmental performance among school age children in rural Guatemala is associated with prenatal and postnatal exposure to carbon monoxide, a marker for exposure to woodsmoke

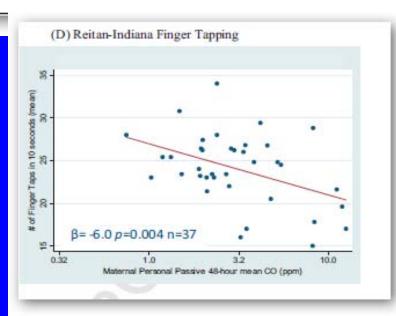
u Linda Dix-Cooper^a, Brenda Eskenazi^b, Carolina Romero^c, John Balmes^{a,d}, Kirk R. Smith^{a,*}

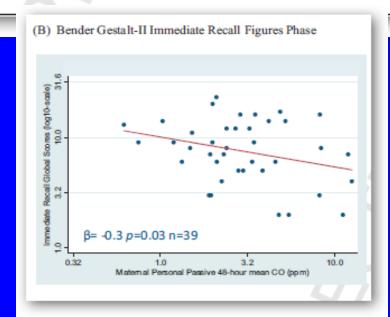
* Division of Environmental Health Sciences, School of Public Health, University of California, Berkeley, CA 94720-7360, USA

^b Center for Environmental Research and Children's Health (CERCH), School of Public Health, University of California, Berkeley, CA, USA

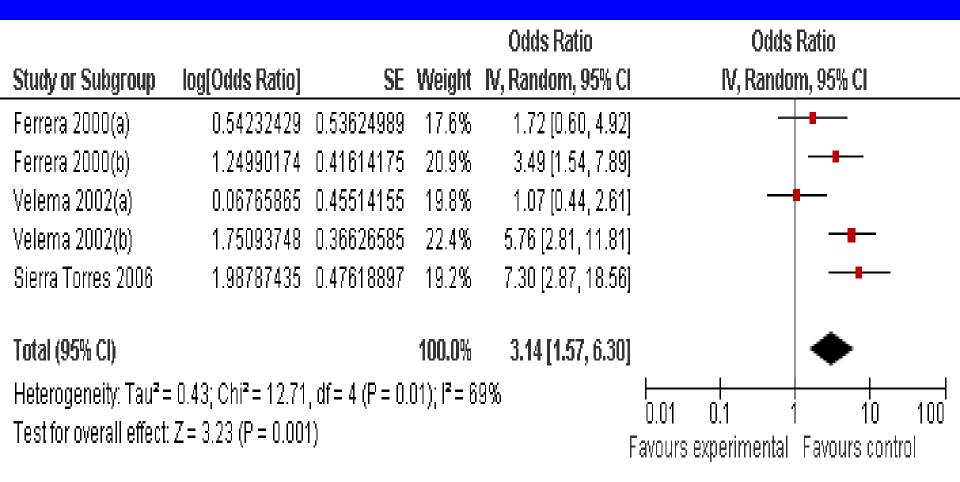
Centro de Estudios en Salud Universidad Del Valle, Guatemala

^d Division of Occupational and Environmental Medicine, Department of Medicine, University of California, San Francisco, CA, USA





Cervical Cancer and Household Air Pollution



Three papers; two done in Honduras, one in Columbia

Issues with the Cervical Cancer Studies.

- Problems
 - Poor exposure assessment
 - Not all adjusted for smoking
 - Doubtful control selection
 - Interaction with HPV not clear
- Strengths
 - Large effects
 - Exposure-response shown
 - Known effects of tobacco smoke

Does a chimney reduce exposure enough?

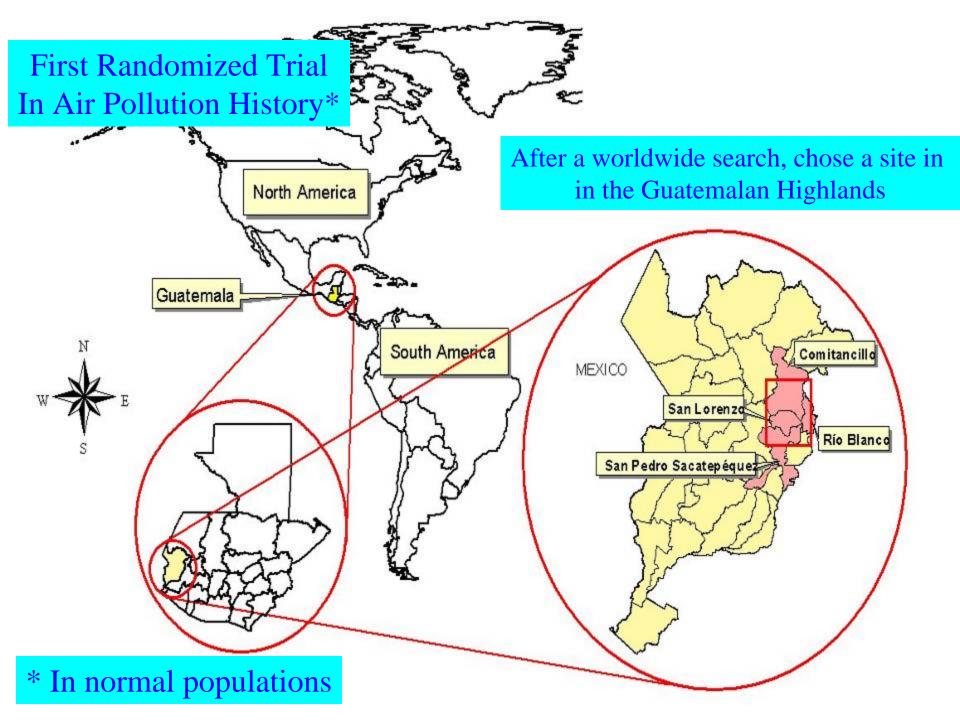
THELANCET-D-09-06268R3 S0140-6736(11)60921-5

Embargo: [add date when known]

Effect of reduction in household air pollution on childhood pneumonia in Guatemala (RESPIRE): a randomised controlled trial

Kirk R Smith, John P McCracken, Martin W Weber, Alan Hubbard, Alisa Jenny, Lisa M Thompson, John Balmes, Anaite Diaz, Byron Arana, Nigel Bruce

Published Nov 2011



RESPIRE – Randomized trial (n=518) Impact on pneumonia up to 18 months of age



Traditional open 3-stone fire: kitchen 48-hour PM_{2.5} levels of 600 - 1200 µg/m³



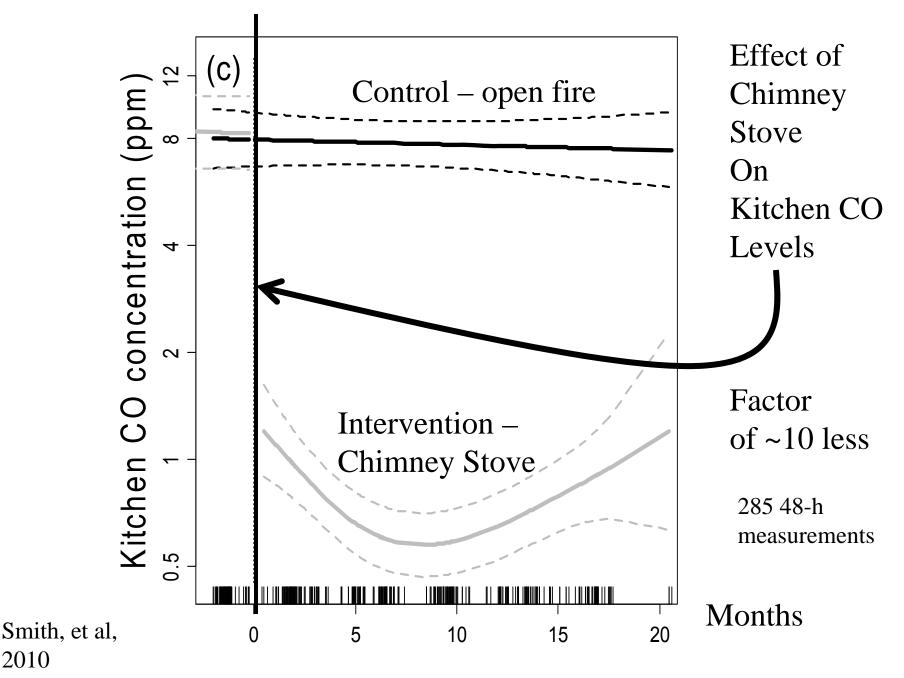
Chimney wood stove, locally made and popular with households

CO monitor

CO monitor

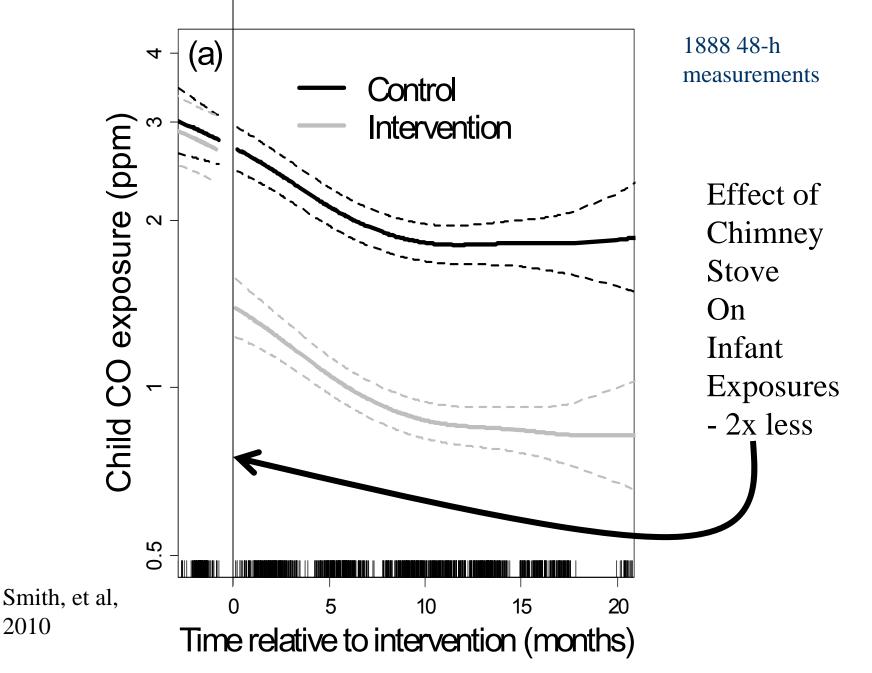
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Guatemala RCT: Kitchen Concentrations



2010

Infant Exposures

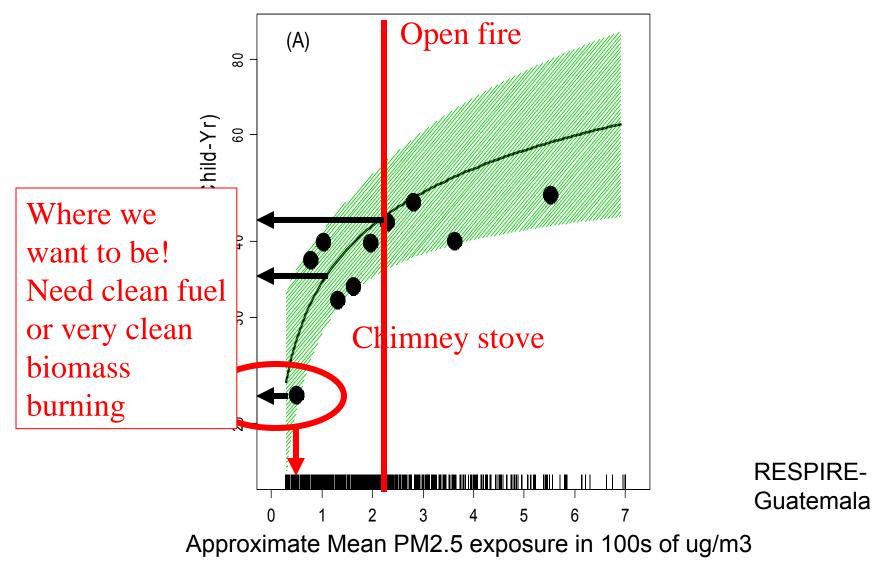


Good chimney stove caused kitchen levels to go down by 10x, but child exposures were down by only 2x, because

- --Time-activity: the kids do not spend their entire day in the kitchen
- --Household (or "neighborhood") pollution: a chimney does not reduce smoke, but just shifts it outside into the household environment, where the difference between intervention and control households was less
 --No significant difference in bedrooms



MD-diagnosed Acute Lower Respiratory Infection



You have heard of secondhand smoke – from tobacco burning

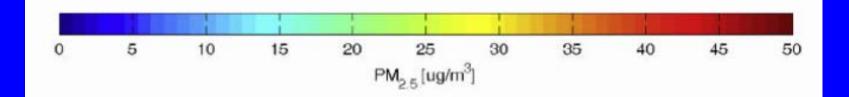
But there is another kind – from cookfires



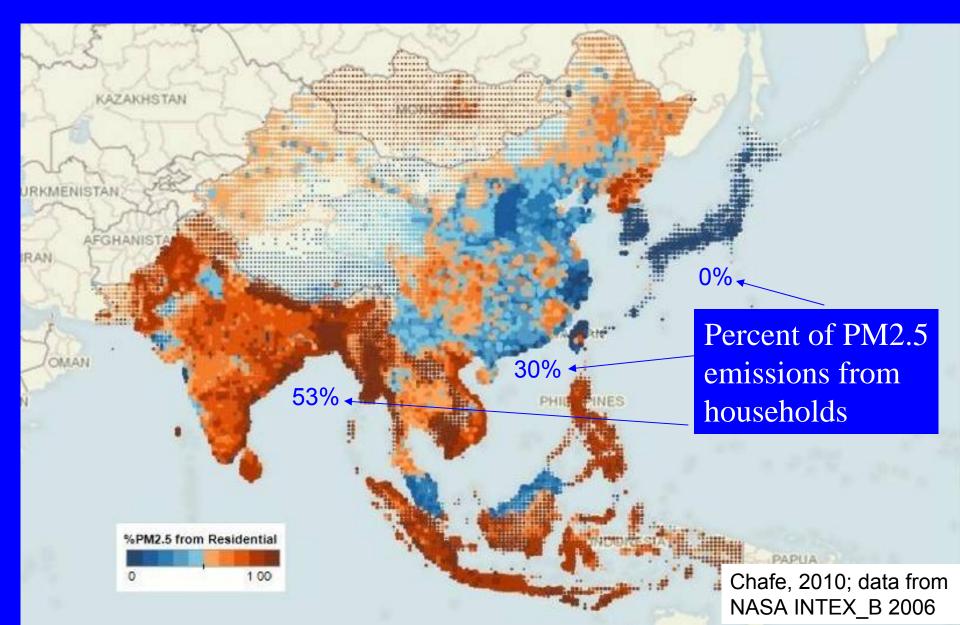
45

MODIS

Large areas of rural India and China have high ambient air pollution – much from household fuel



Percent PM_{2.5} emissions from households



Combustion Particles

The Generalized Exposure Response (GER)

Heart Disease and Combustion Particle Doses

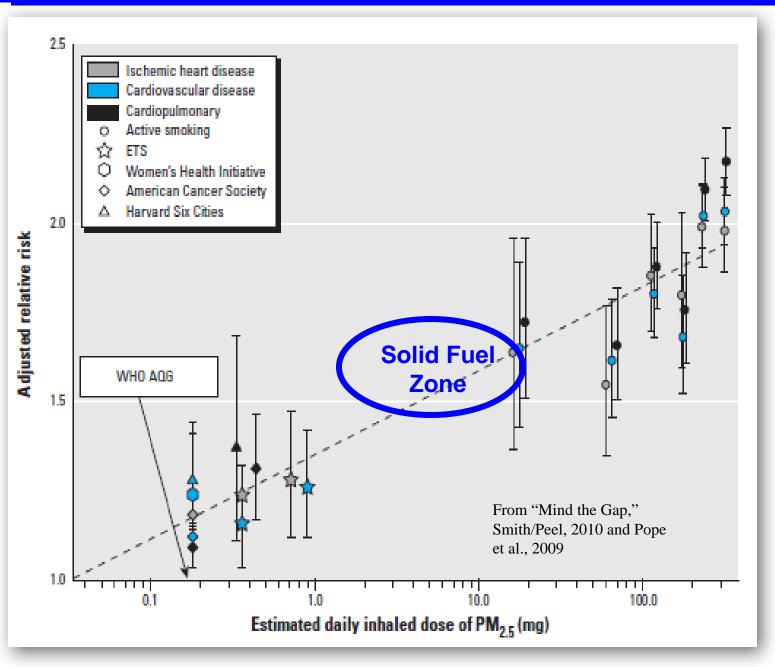
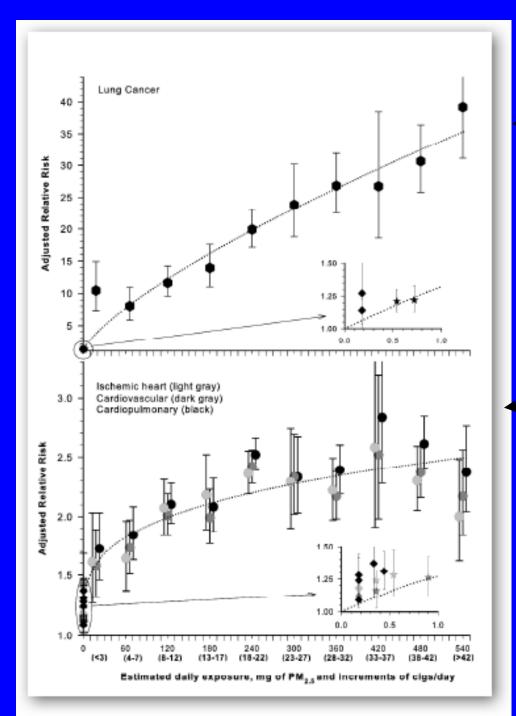


Table 2. Adjusted relative risk estimates^a for various increments of exposure from cigarette smoking (versus never smokers), second hand cigarette smoke, and ambient air pollution from the present analysis and selected comparison studies.

Source of risk estimate	Increments of Exposure	Adjusted RR (95% CI)				Estimated Daily
		Lung Cancer	IHD	CVD	CPD	Dose PM2.5 (mg)b
ACS- present analysis	<3 (1.5) cigs/day	10.44 (7.30-14.94)	1.61 (1.27-2.03)	1.58 (1.32-1.89)	1.72 (1.46-2.03)	18
ACS- present analysis	4-7 (5.5) cigs/day	8.03 (5.89-10.96)	1.64 (1.37-1.96)	1.73 (1.51-1.97)	1.84 (1.63-2.08)	66
ACS- present analysis	8-12 (10) cigs/day	11.63 (9.51-14.24)	2.07 (1.84-2.31)	2.01 (1.84-2.19)	2.10 (1.94-2.28)	120
ACS- present analysis	13-17 (15) cigs/day	13.93 (11.04-17.58)	2.18 (1.89-2.52)	1.99 (1.77-2.23)	2.08 (1.87-2.32)	180
ACS- present analysis	18-22 (20) cigs/day	19.88 (17.14-23.06)	2.36 (2.19-2.55)	2.42 (2.28-2.56)	2.52 (2.39-2.66)	240
ACS- present analysis	23-27 (25) cigs/day	23.82 (18.80-30.18)	2.29 (1.91-2.75)	2.33 (2.02-2.69)	2.33 (2.03-2.67)	300
ACS- present analysis	28-32 (30) cigs/day	26.82 (22.54-31.91)	2.22 (1.97-2.49)	2.17 (1.98-2.38)	2.39 (2.19-2.60)	360
ACS- present analysis	33-37 (35) cigs/day	26.72 (18.58-38.44)	2.58 (1.91-3.47)	2.52 (1.98-3.19)	2.83 (2.28-3.52)	420
ACS- present analysis	38-42 (40) cigs/day	30.63 (25.79-36.38)	2.30 (2.05-2.59)	2.37 (2.16-2.59)	2.61 (2.40-2.84)	480
ACS- present analysis	43+ (45) cigs/day	39.16 (31.13-49.26)	2.00 (1.62-2.48)	2.17 (1.84-2.56)	2.37 (2.04-2.76)	540
ACS-air pol. original	24.5 µg/m ³ ambient PM2.5			-tota	1.31(1.17-1.46)	0.44
ACS-air pol. extend.	10 µg/m ³ ambient PM2.1	1.14(1.04-1.23)	1.18(1.14-1.23)	1.12(1.08-1.15)	1.09(1.03-1.16)	0.18
HSC-air pol. original	18.6 µg/m3 ambient PM25		and an an an and a start of the		1.37(1.11-1.68)	0.33
HSC-air pol. extend.	10 µg/m ³ ambient PM _{2.5}	1.21(0.92-1.69)		1.28(1.13-1.44)		0.18
WHI-air pol.	10 µg/m ³ ambient PM _{2.5}		12/194	1.24(1.09-1.41) ^c	0.000	0.18
SGR-SHS	Low- moderate SHS exp.			1.16(1.03-1.32)		0.36
SGR-SHS	Moderate-high SHS exp			1.26(1.12-1.42)		0.90
SGR-SHS	Live with smoking spouse	1.21(1.13-1.30)				0.54
SGR-SHS	Work with SHS exposure	1.22(1.13-1.33)				0.72
INTERHEART	1-7 hrs/wk SHS exp.		1.24(1.17-1.32) ^d			0.36
INTERHEART	Live with smoking spouse		1.28(1.12-1.47) ^d			0.54

Pope et al. Environmental Health Perspectives 2011, in press



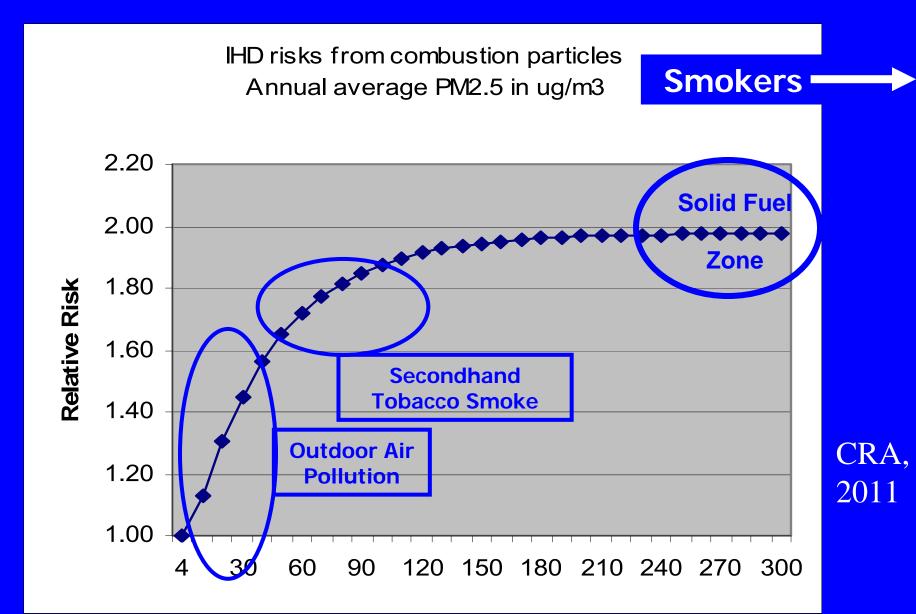
Heart Disease

Lung

Cancer

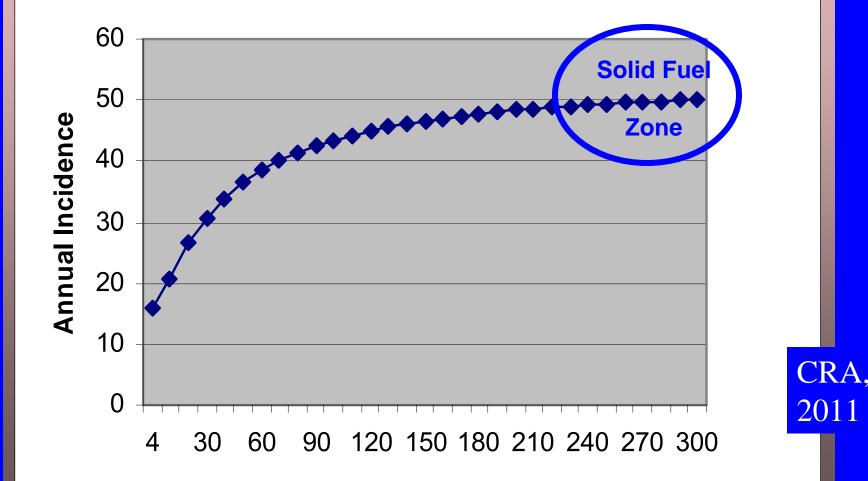
Pope et al. Environmental Health Perspectives 2011

Generalized Exposure-Response: Outdoor Air, SHS, and Smoking



Generalized Exposure-Response: Outdoor Air, SHS, and HAP

Pneumonia from combustion particles Annual average PM2.5 in ug/m3



What we can expect based on ATS epidemiological evidence?

- "Traditional" ATS cancers: lung, oral cavity, pharynx, larynx, oesophagus, pancreas, urinary bladder and renal pelvis
- "New" ATS cancers: nasal, sinus, nasopharynx, stomach, liver, kidney, uterine cervix, oesophagus, and leukaemia

Review of Epi Evidence: Lung Cancer, 2004

ETS/SHS and Cancer?

- Causes lung cancer in nonsmoking adults.
 ...living with a smoker increases a nonsmoker's chances of developing lung cancer by 20 to 30 percent.
- Some research suggests that SHS/ETS may increase the risk of breast cancer, nasal sinus cavity cancer, and nasopharyngeal cancer in adults and the risk of leukemia, lymphoma, and brain tumors in children.

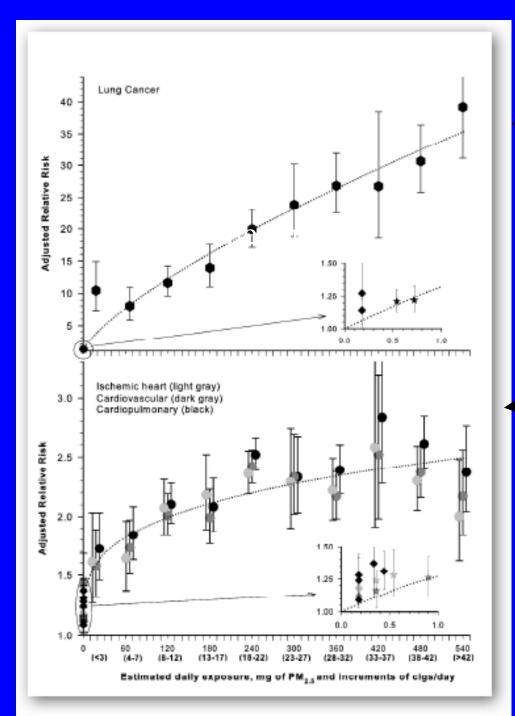
NCI Factsheet, 2012

The cancer research challenge

- Hypothesis #1: Most if not all the same cancers found for ATS can also be found for ETS/SHS and HAP, albeit at appropriately lower risks
- Hypothesis #2: They have not yet all been confirmed for ETS/SHS because lower exposures and resultant risk levels make detection difficult compared to ATS studies.
- Living with substantially higher exposures than SHS/ETS, even if lower than ATS, HAP-exposed populations offer excellent opportunities for pinning down a range of cancer risks and associated mutagenicity and genetic associations
- Considering that the HAP-population is larger and younger than those for ATS or ETS/SHS, the total cancer global burden of cancer is likely to be substantial.

Two policy dilemmas

- How to think about protection when some major impacts are linear, e.g. lung cancer, and others highly non-linear, e.g., heart disease?
- How to think about protection when the major reductions in risk occur only at relatively low levels, or put differently, there is relatively little benefit by changing from high to moderate levels of exposure?



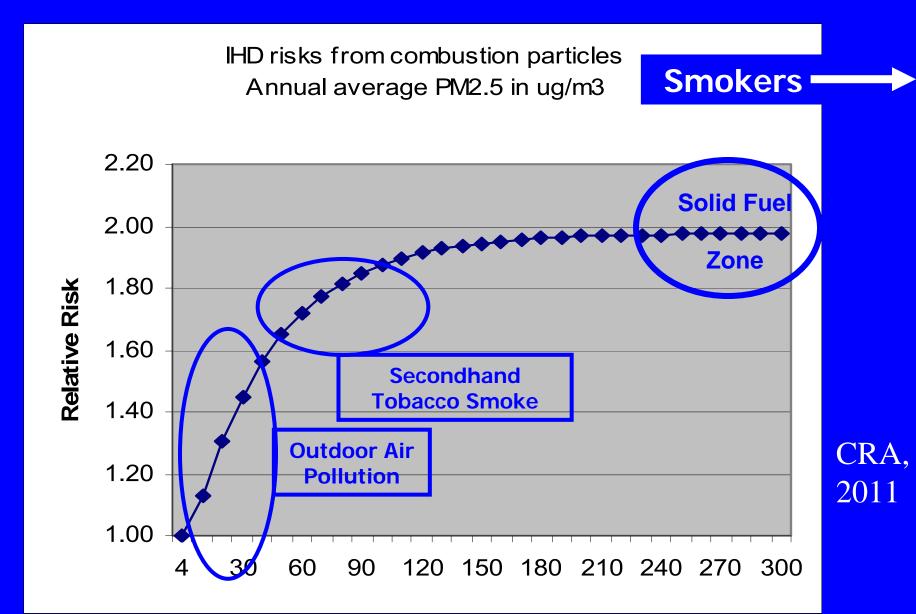
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Pope et al. <u>Environmental</u> <u>Health</u> <u>Perspectives</u> 2011, in press

Generalized Exposure-Response: Outdoor Air, SHS, and Smoking



Many thanks

Publications and presentations on website – easiest to just "google" Kirk R. Smith

