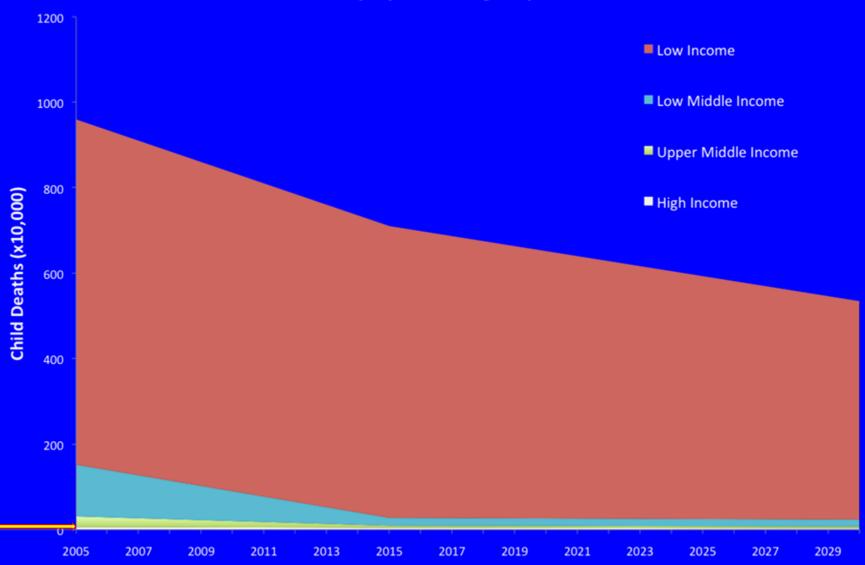
Indoor Air Pollution and Child Health What do we know, what should we know and what are some of the systems relationships?

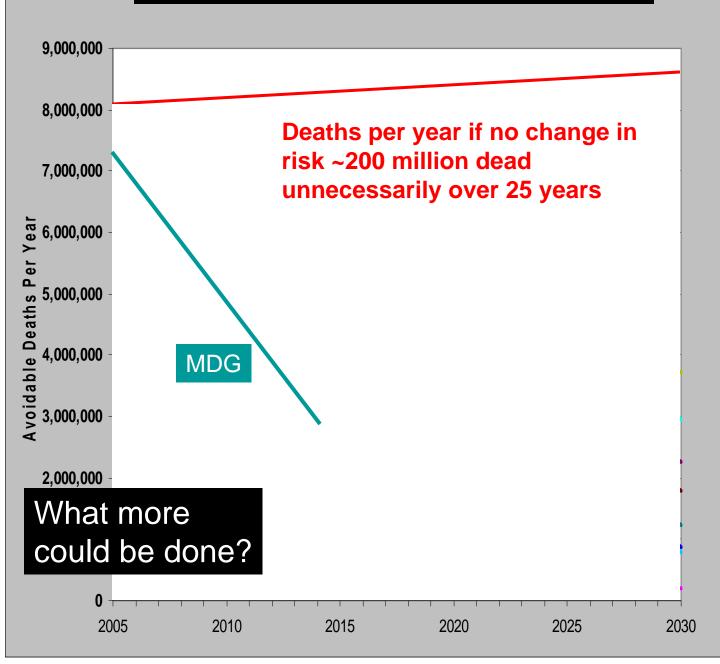
> Kirk R. Smith, PhD, MPH Professor of Global Environmental Health University of California, Berkeley

> > NICHD/NIEHS, Bethesda September 26, 2008

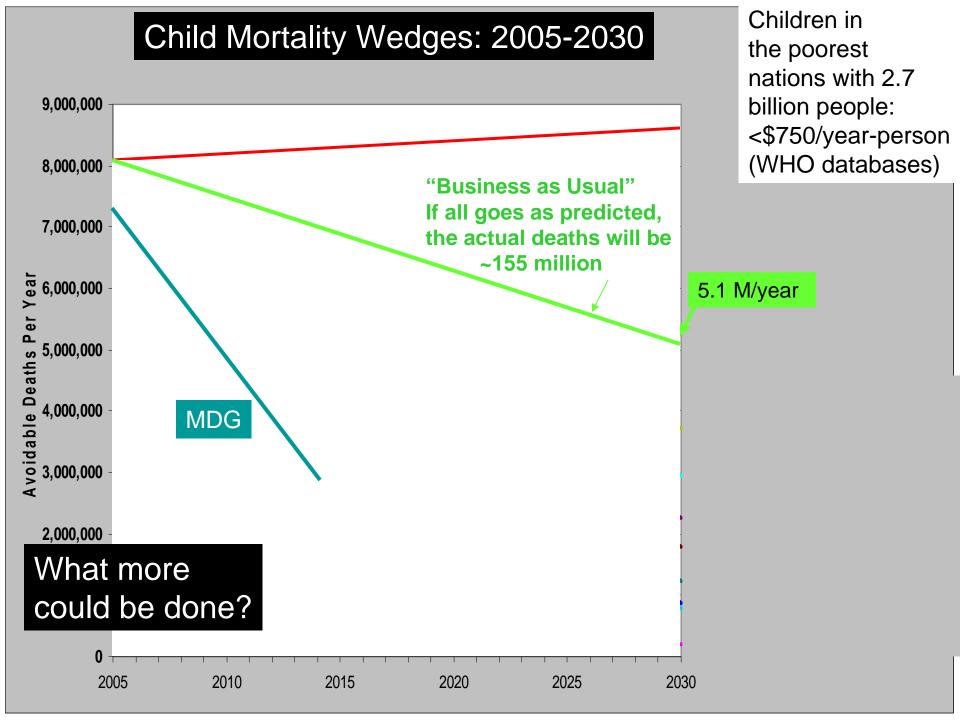


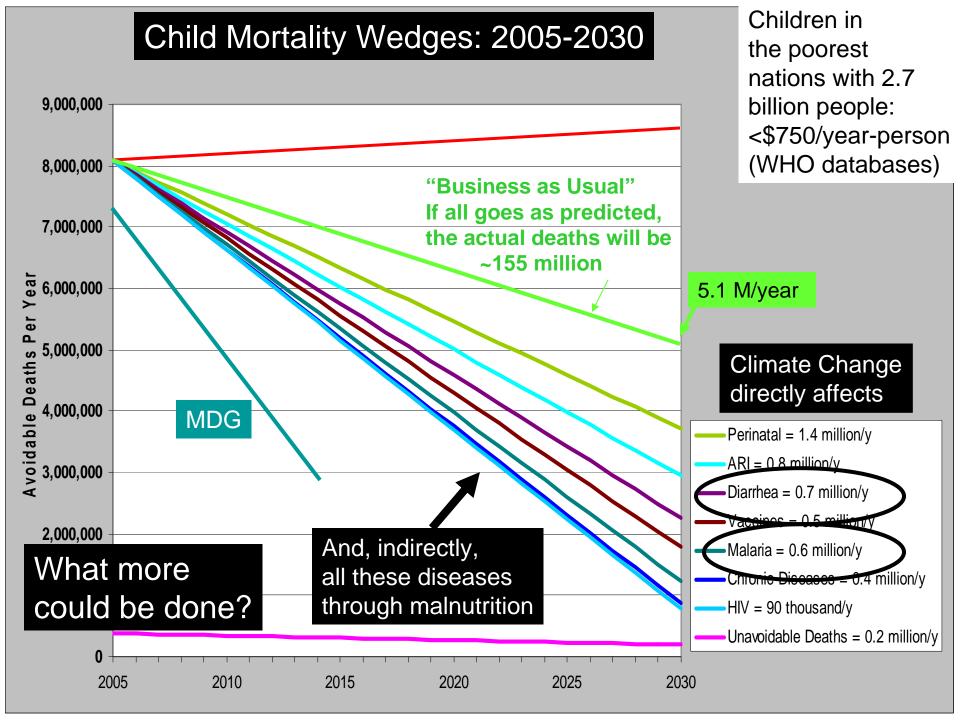
0-4 mortality by income group 2005-2030

Child Mortality Wedges: 2005-2030

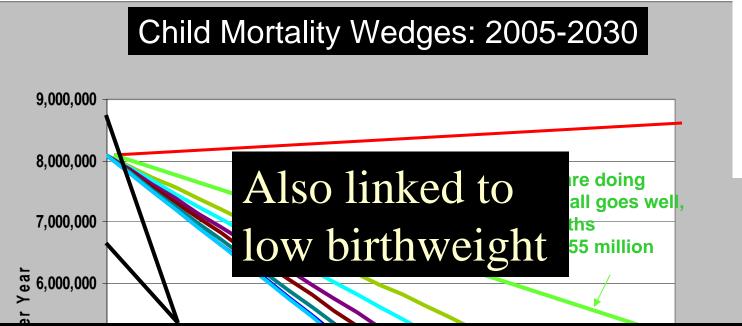


Children in the poorest nations with 2.7 billion people: <\$750/year-person (WHO databases)



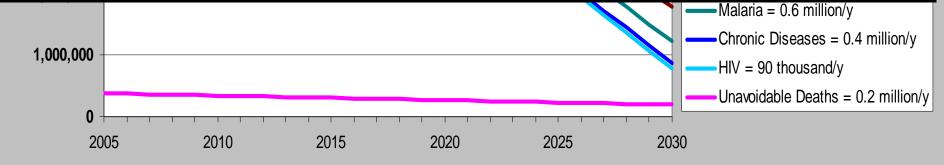


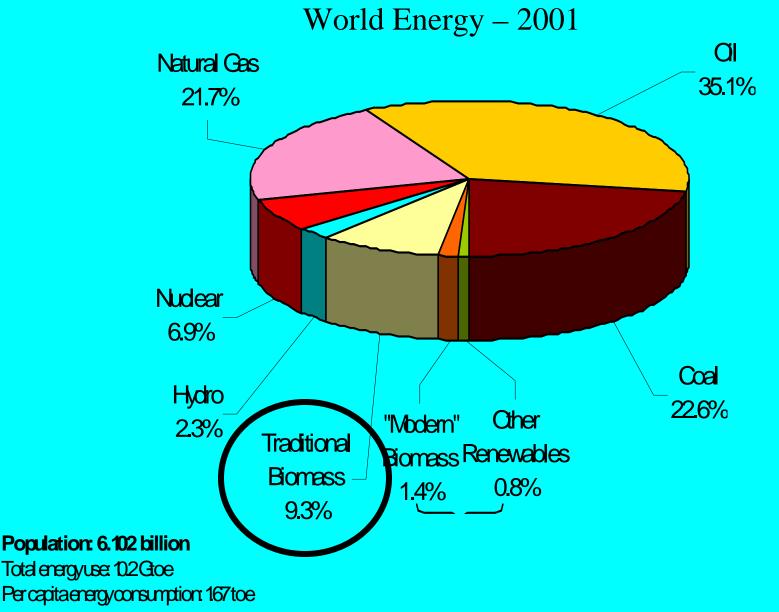
Children in Child Mortality Wedges: 2005-2030 the poorest nations with 2.7 9,000,000 billion people: <\$750/year-person (WHO databases) 8,000,000 However, we are doing something. If all goes well, 7,000,000 the actual deaths will be ~155 million Year 6,000,000 er Δ. 2+ million of these 5,000,000 Avoidable Deaths child deaths are due to Acute Respiratory 4,000,000 Infections – ARI Perinatal = 1.4 million/y These kill more children ARI = 0.8 million/y3,000,000 Diarrhea = 0.7 million/y than any other condition today Vaccines = 0.5 million/y 2,000,000 Malaria = 0.6 million/y Chronic Diseases = 0.4 million/y 1,000,000 HIV = 90 thousand/y Unavoidable Deaths = 0.2 million/y 0 2005 2010 2015 2020 2025 2030



Children in the poorest nations with 2.7 billion people: <\$750/year-person (WHO databases)

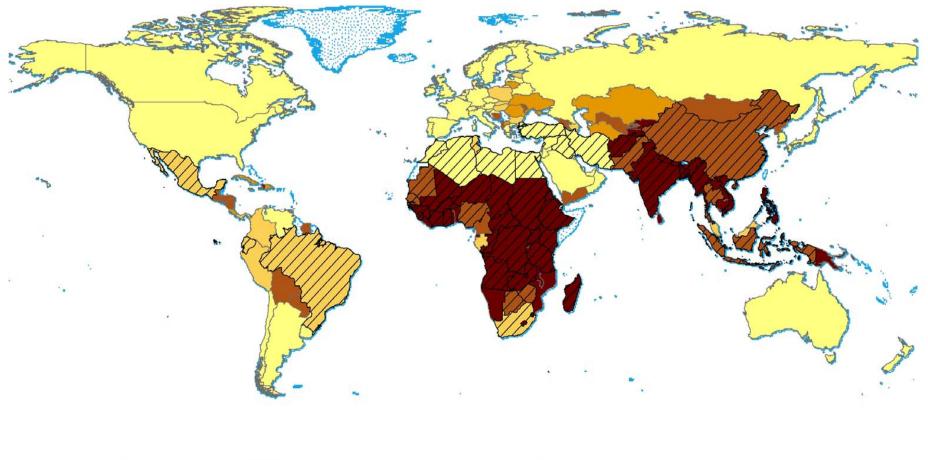
ARI has strong links to inefficient energy use, which offers a great opportunity for "Co-benefits" -- achieving both climate mitigation and health protection with the same policies

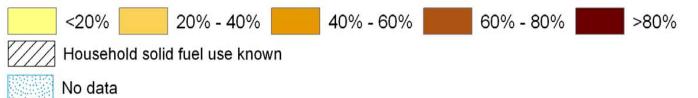


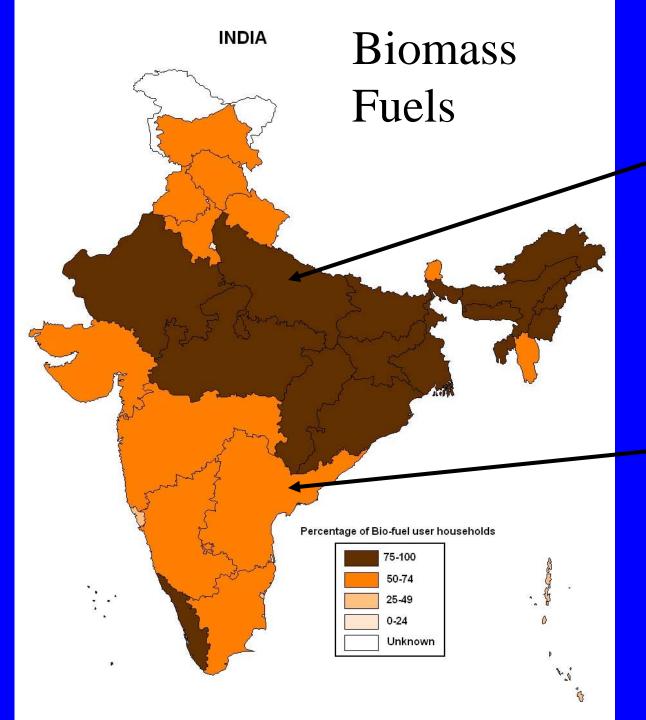


World Energy Assessment, 2004

National Household Solid Fuel Use, 2000







More than 75% of households

50-74% of households

2000 Census

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)?

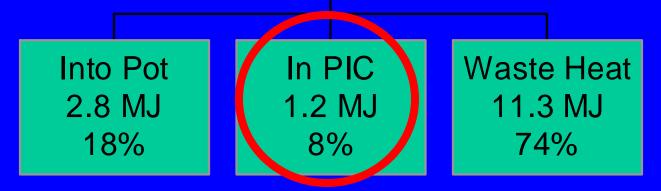


Reason: the combustion efficiency is far less than 100%

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



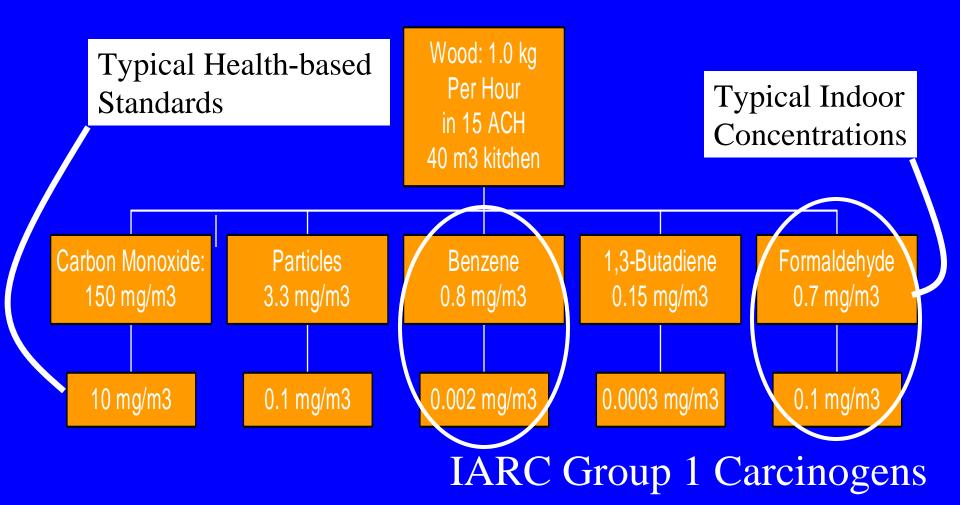
PIC = products of incomplete combustion = CO, HC, C, etc.

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 Air Pollutants From Typical Woodfired Cookstove in India.



First person in human history to have her exposure measured doing one of the oldest tasks in human history

Exposures seem to be high in a large vulnerable population. But what are the health effects?

Kheda District Gujarat, India 1981 <u>ALRI/</u> <u>Pneumonia</u> (meningitis)

Low birth weight & stillbirth

Asthma

Early infant Death?

> Cognitive Effects?

Diseases for which we have epidemiological studies linking indoor air pollution to disease <u>Chronic</u> <u>obstructive</u> <u>lung disease</u>

Interstitial LD

<u>Cancer</u> (lung, NP, cervical, aero-digestive)

Blindness (cataracts, trachoma)

Tuberculosis

Heart disease?

Study design	Ν	OR	95% CI		
	*				
Intervention	2	1.28	1.06, 1.54		
New ALRI-IAP					
Systematic					
Bystematic					

Review and Meta-Analysis

Dherani et al. Bull WHO, 2008

Cross- sectional	3	1.49	1.21, 1.85
All	26	1.78	1.45, 2.18

Study or sub-category	Odds Ratio (random) 95% Cl	VVeight %	Oc	lds Ratio (95%	
01 Intervention Studies					
Smith(2007)a		5.53	1.18	[0.88,	1.58]
Smith(2007)b		5.73	1.35	[1.05,	1.73]
Subtotal (95% CI)	◆	11.26	1.28	[1.06,	1.54]
Test for heterogeneity: Chi ² = 0.4	48, df = 1 (P = 0.49), ² = 0%				
Test for overall effect: Z = 2.54 ((P = 0.01)				
02 Cohort Studies					
Armstrong(1991)a		2.80	0.50	[0.20,	1.22]
Armstrong(1991)b		3.65	1.90	[0.96,	3.75]
Cambell(1989)	· · · · · · · · · · · · · · · · · · ·	3.25	2.80	[1.29,	6.08]
Ezzati(2001)		3.86	2.33	[1.23,	4.40]
Jin(1993)		5.69	0.80	[0.62,	1.03]
Pandey(1989)a		4.34	2.45	[1.43,	4.19]
Pandey(1989)b		1.52	40.65	[9.79,	168.75]
Subtotal (95% CI)		25.11	2.12	[1.05,	4.25]
Test for heterogeneity: Chi ² = 54 Test for overall effect: Z = 2.11 (.07, df = 6 (P < 0.00001), l² = 88.9% (P = 0.03)				
03 Case-Control Studies					
Azizi(1995)		3.97	1.20	[0.65,	2.21]
Broor(2001)		4.49		[1.51,	
Collings(1990)		4.85	2.16	[1.40,	3.33]
De Francisco(1993)		2.15		[1.72,	
Fonsecca(1996)		4.68		[0.71,	
Johnson(1992)a		3.15	0.80	[0.36,	1.78]
Kossove(1982)		→ 1.96	4.77	[1.44,	15.74]
Kumar(2004)	· · · · · · · · · · · · · · · · · · ·	2.45	3.87	[1.42,	10.57]
Mahalanabas(2002)	2	— 3.63	3.97	[2.00,	7.88]
Morris(1990)			4.85	[1.75,	13.40]
O'Dempsey(1996)		- 2.59	2.55	[0.98,	6.64]
Robin(1996)a		2.95	1.40	[0.60,	3.28]
Victora(1994)a		4.08		[0.61,	
Wayse(2004)		2.90		[0.58,	
Wesley(1996)		1.87		[0.39,	
Subtotal (95% CI)	••••••••••••••••••••••••••••••••••••••	48.15	1.97	[1.47,	2.64]
Test for heterogeneity: Chi ² = 32 Test for overall effect: Z = 4.53 (.72, df = 14 (P = 0.003), l² = 57.2% (P < 0.00001)				
04 Cross-sectional Studies					
Mishra(2003)		3.83	2 20	[1.16,	4 181
Mishra(2005)		5.87		[1.28,	
Wichmann(2006)		5.79	1.29	[1.02,	1.631
Subtotal (95% Cl)		15.48	1.49	[1.21,	1.851
Test for heterogeneity: Chi ² = 3.1	19. df = 2 (P = 0.20), ² = 37.3%				
Test for overall effect: Z = 3.74 (
Total (95% Cl)	•	100.00	1.78	[1.45,	2.18]
	1.74, df = 26 (P < 0.0001), l² = 74.4%	6			
Test for overall effect: Z = 5.61 ((P < 0.00001)	1			
0.	1 0.2 0.5 1 2 5	10			
	Increased risk Decreased risk	6			

History of a RCT

- ~1980: early studies of health effects in Nepal and elsewhere
- 1981: first measurements of pollution levels in India
- 1984: international meeting to decide on needed research
 - Chose randomized control trial (RCT) of ALRI
- 1986-89: unfunded proposals to do RCT in Nepal
- 1990: WHO establishes committee to find best sites
- 1992: Guatemala chosen
- 1991-1999: Pilot studies to establish data needed for proposal
- 1996-1999: unfunded proposals
- 2001: NIH funds first randomized control trial for air pollution in highland Guatemala
- 2002-2005: fieldwork completed
- 2007: first results published
- 23 years from deciding to conduct RCT to results! RESPIRE - Guatemala

First Randomized Control Trial in Air Pollution History



Traditional 3-stone open fire

Plancha chimney wood stove

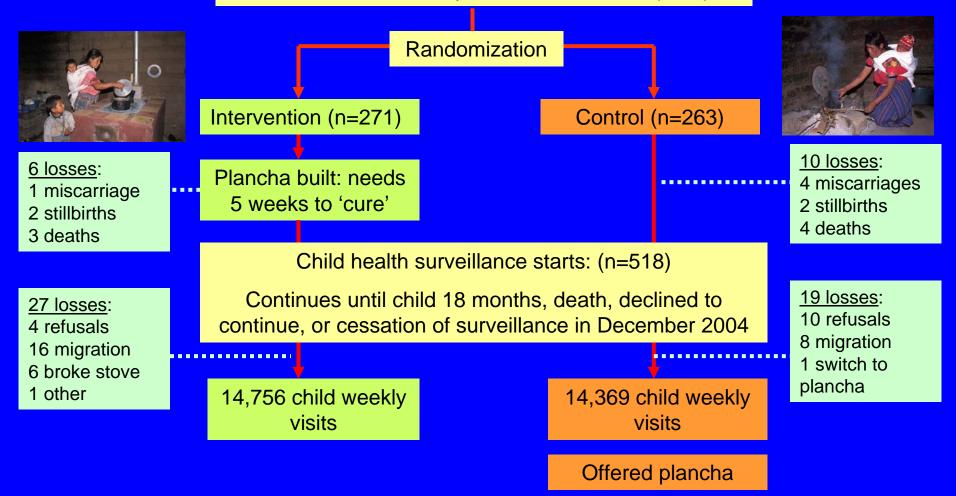
RESPIRE - Guatemala

RCT Study flow chart

October 2002 - Eligible homes: open fire; pregnant; child <4 months: n=777

> Declined n=243 (31%)

Consent & baseline questionnaire: n=534 (69%)



Guatemala RCT Results

[Unpublished results presented] showing that with ITT analysis, all outcomes showed improvement, but only severe ALRI outcomes were significant. With exposureresponse analysis using personal pollution monitoring, however, nearly all outcomes significant, including primary outcome: MD-diagnosed pneumonia] **Respiratory Disease - other**

- Minor burden, but important economic impact
 - Otitis Media 0.08% of global burden
 - AURI 0.11%
- Compared to 3% and 6% for LBW and ALRI

Low birth weight (associated with IAP)

- Large direct impact on child mortality
- Of which perhaps 25% is ALRI
- Barker-type effect has been found on chronic disease over life time, including COPD

Low Birth Weight

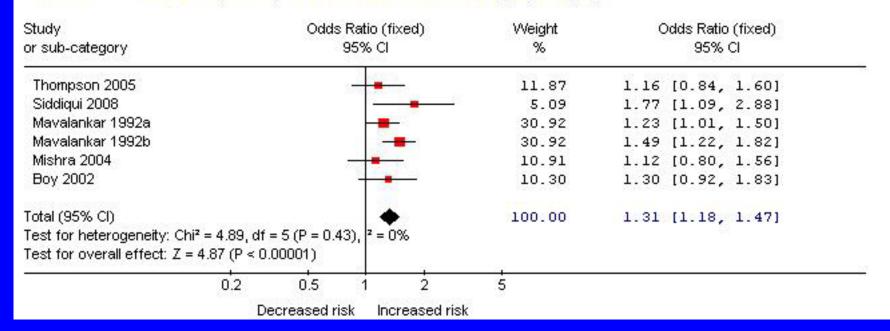
- Small number of studies conducted appearing to show results for IAP consistent with evidence from ETS and ambient air pollution.
- Pregnant women in countries with high rates of (a) solid fuel use, and (b) adverse pregnancy outcomes rarely able to avoid activities that expose them and their unborn children to IAP
- Hence, even modest increase in RR for these conditions can be expected to translate into a substantial population attributable risk.
- Systematic review carried out in order to:
 - summarise the extent and quality of the evidence for the relationship between IAP from solid fuel use and adverse pregnancy outcomes
 - quantify these associations by meta-analysis.

Results: IAP and LBW (<2500gms)

 Review:
 Indoor air pollution from solid fuels and risk of low birth weight and stillbirth: a systematic review and meta-analysis.

 Comparison:
 01 Systematic review of indoor air pollution and low birth weight

 Outcome:
 01 Meta-analysis : Exposure to IAP and % Low Birth Weight (<2500gms)</td>



Heterogeneity - $(I^2 = 0\%; Chi^2 (df=5) = 4.89, p=0.43)$ Funnel plot asymmetry - Begg's test (p = 0.566) Egger's test (p=0.989)

Pope et al., in prep

Results: IAP and birth weight

Review:	Indoor air pollution from solid fuels and risk of low birth weight and stillbirth: a systematic review and meta-analysis.			
Comparison:	03 Systematic review of indoor air pollution and low birth weight			
Outcome:	01 Meta-analysis: Exposure to IAP and mean birth weight			
Study	WMD (fixed)	Weight	VVMD (fixed)	
or sub-categor	y 95% Cl	%	95% CI	
Thompson 20	05	14.89	47.00 [-69.69, 163.69]	
Siddiqui 2008		24.66	82.00 [-8.69, 172.69]	
Mishra 2004		12.98	175.00 [50.00, 300.00]	
Boy 2002		47.47	63.00 [-2.36, 128.36]	
Total (95% CI)	•	100.00	79.84 [34.81, 124.87]	
Test for hetero	geneity: Chi² = 2.79, df = 3 (P = 0.43), ² = 0%			
Test for overall	l effect: Z = 3.47 (P = 0.0005)	3		
	-1000 -500 0 500	1000	2	
	Not open fire Open fire			

Heterogeneity - (I² = 0%; Chi² (df=3) = 2.79, p=0.43)

Funnel plot asymmetry - Begg's test (p = 0.308) Egger's test (p=0.479)

Pope et al., in prep

Results: IAP and still birth

 Review:
 Indoor air pollution from solid fuels and risk of low birth weight and stillbirth: a systematic review and meta-analysis.

 Comparison:
 05 Systematic review of indoor air pollution and stillbirth

 Outcome:
 01 Meta-analysis: Exposure to IAP and stillbirth

Study or sub-category	Odds Ratio (fixed) 95% Cl	VVeight %	Odds Ratio (fixed) 95% Cl	
Siddiqui 2005		- 17.29	1.90 [1.12, 3.23]	335
Mavalankar 1991		35.28	1.50 [1.04, 2.17]	
Mishra 2005		47.43	1.44 [1.05, 1.98]	
Total (95% Cl)	•	100.00	1.53 [1.23, 1.91]	
Test for heterogeneity: Chi ² = I	0.79, df = 2 (P = 0.67), ² = 0%			
Test for overall effect: Z = 3.8	이 2010년 2011년 2011년 1월 18일			
	0.2 0.5 1 2	5		
	Decreased risk Increased ris	sk		

Heterogeneity - (I² = 0%; Chi² (df=2) = 0.79, p=0.67)

Funnel plot asymmetry - Begg's test (p = 0.296) Egger's test (p=0.083)

Pope et al., in prep

Discussion

- Paucity of studies, conducted using a variety of study designs and across a range of settings.
- Despite this, results remarkably consistent with little evidence of statistical heterogeneity.
- Possibility of residual confounding.
- Variation in exposure measurement (need direct measurement for exposure-response analysis).
- Lack of information on gestational age in some of LBW studies.
- Results consistent with literature on ETS, ambient air pollution and LBW.
- Mechanisms: (i) CO (ii) PM
- Few studies on ETS/ ambient air pollution and still birth.

Conclusions

• Population Attributable Risks (%):

LBW = 17.8% Still birth = 27.1% (70% prevalence of solid fuel use)

- Primary or secondary outcome for CRA?
- Strong case for additional studies.

Interactions with IAP

- Arsenic in water "largest poisoning in human history"
 - Causes COPD, lung cancer, and ALRI
 - Much common exposure with IAP
- High occupational exposures to various dusts including silica, asbestos,
- Malnutrition
- Smoking

Large Global Exposures to some surprising pollutants – perhaps largest

- Ultrafine particles fresh and combustiongenerated
- Formaldehyde
- Benzene
- PAH
- Dioxin
- Etc.

Biomass smoke – a global concern

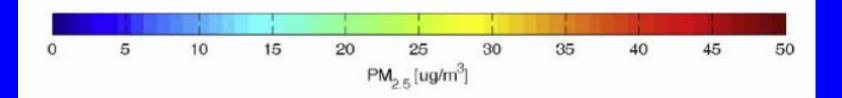
- A significant contribution to PM2.5 emissions around the world – more than half in many developed countries (Canada, Denmark, much of USA, etc.)
 - Ag burning a function of ag production, not income California
 - Wood heating and fireplace use common in many developed countries – Silicon Valley
- Growing because of energy prices
- And climate change
- Not clear whether effects across all major health outcomes are the same as those found in urban studies of PM
 - Chronic and Acute Respiratory
 - Cardiovascular
 - Cancer
- Households in LDCs perhaps only widespread exposure to nearly pure biomass smoke

20-month average ground-level PM2.5 from satellite data

MODIS

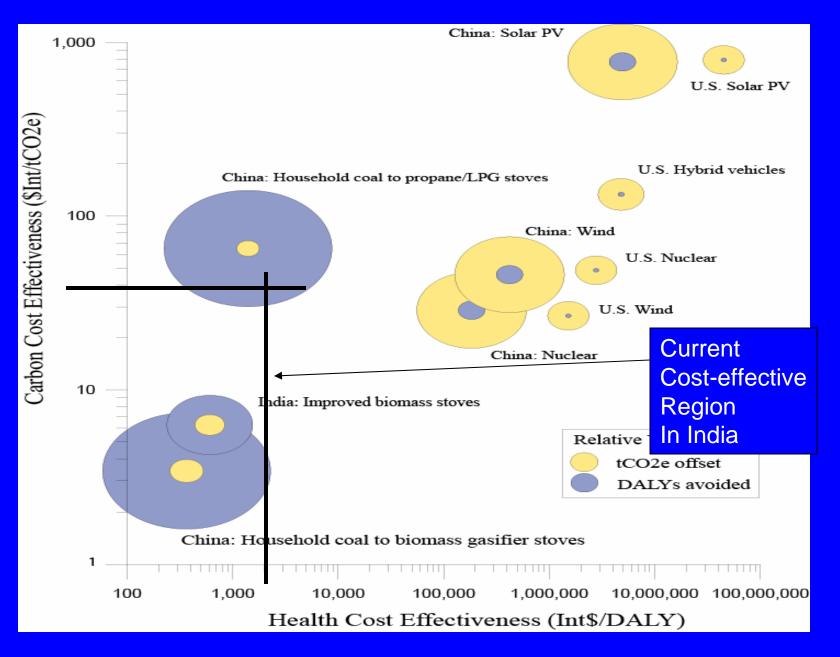
45

Large areas of rural India have high ambient air pollution



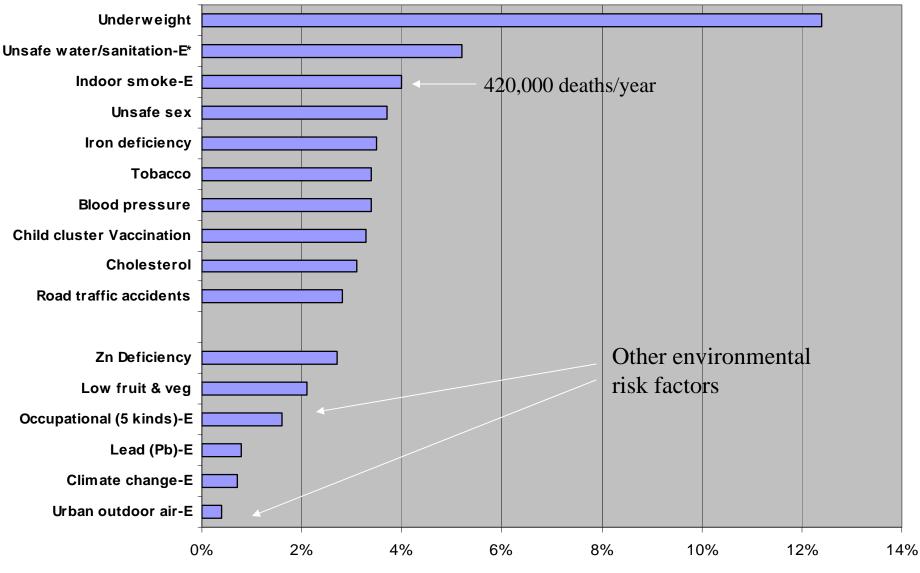
Climate Change – Co-benefits

- Products of incomplete combustion are both health damaging and climate warming
- Efforts to reduce them in household combustion can be highly effective for both goals – co-benefits

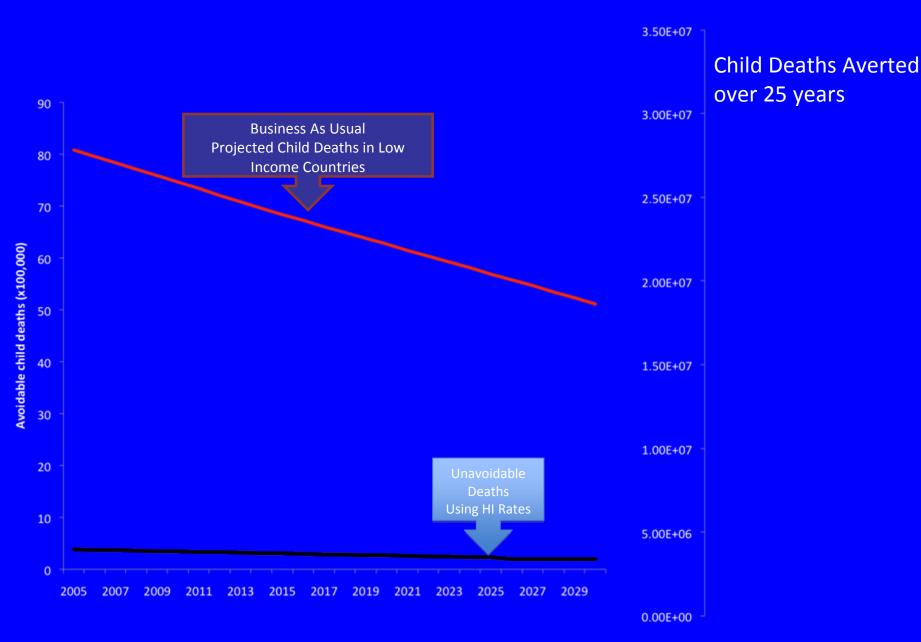


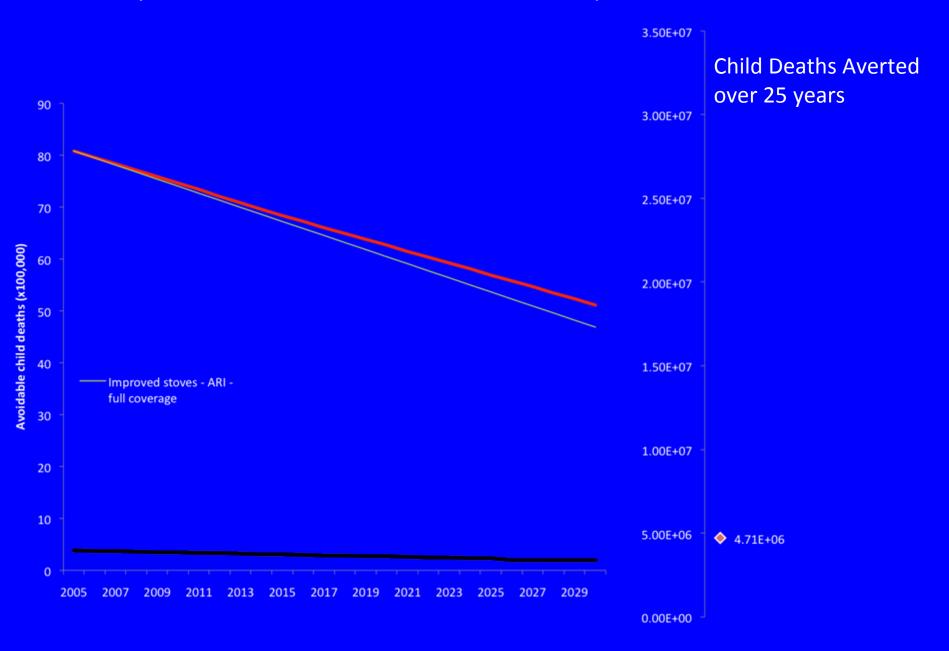
Smith & Haigler, 2008

Indian Burden of Disease from Top 10 Risk Factors and Selected Other Risk Factors

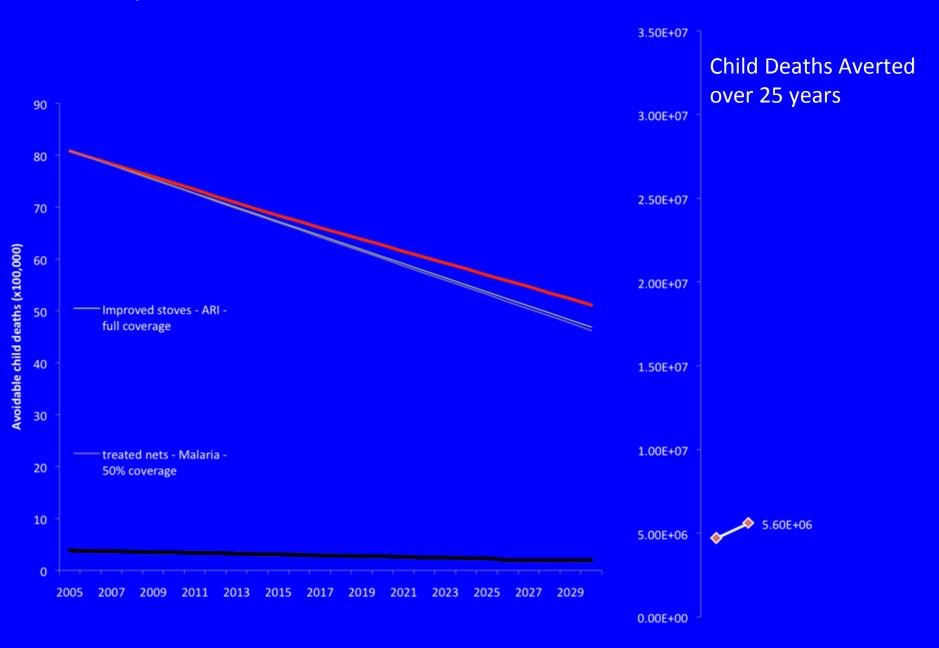


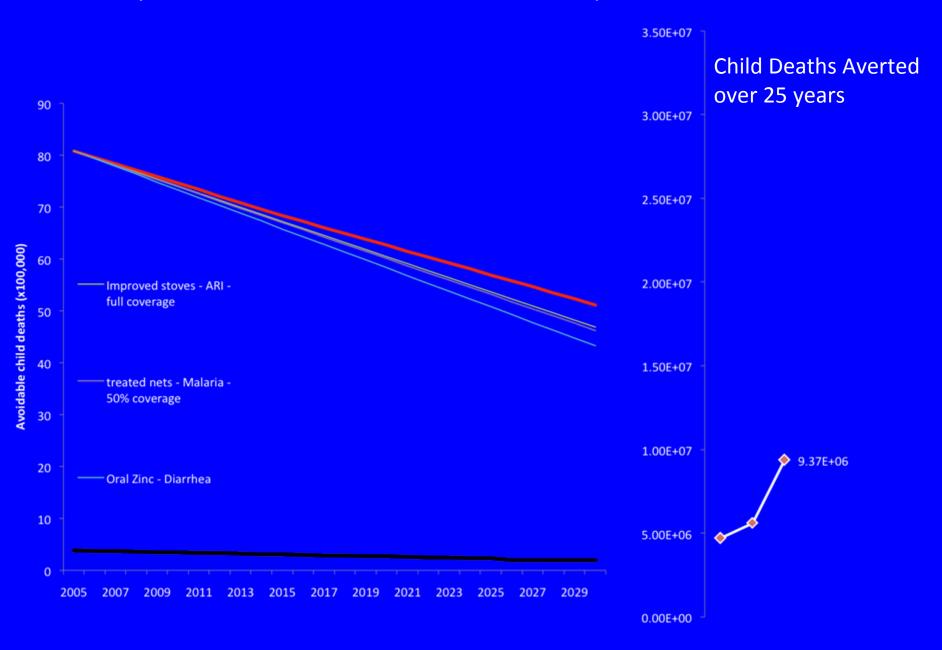
Percent of All DALYs in 2000

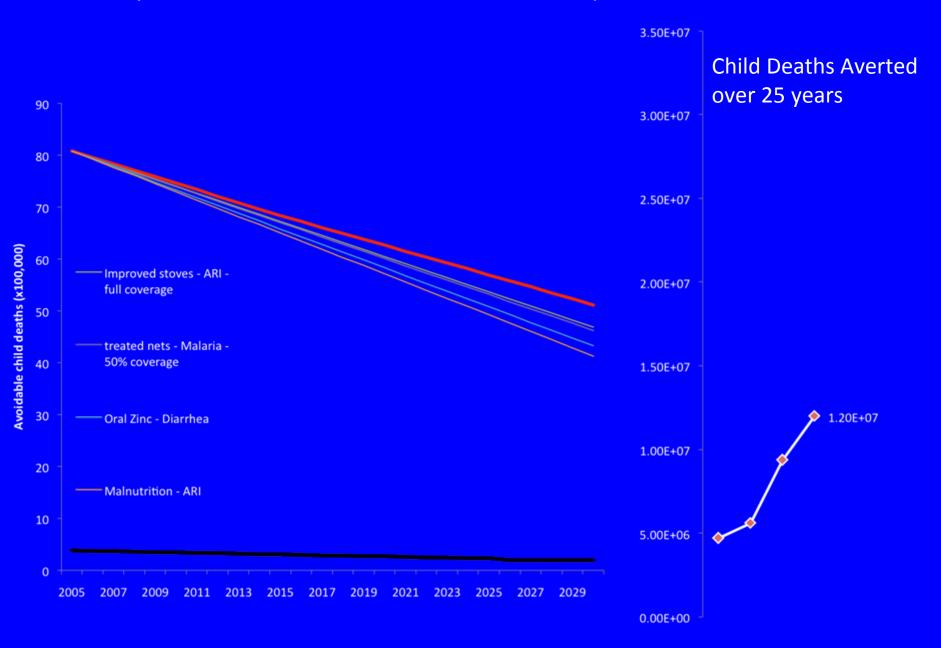


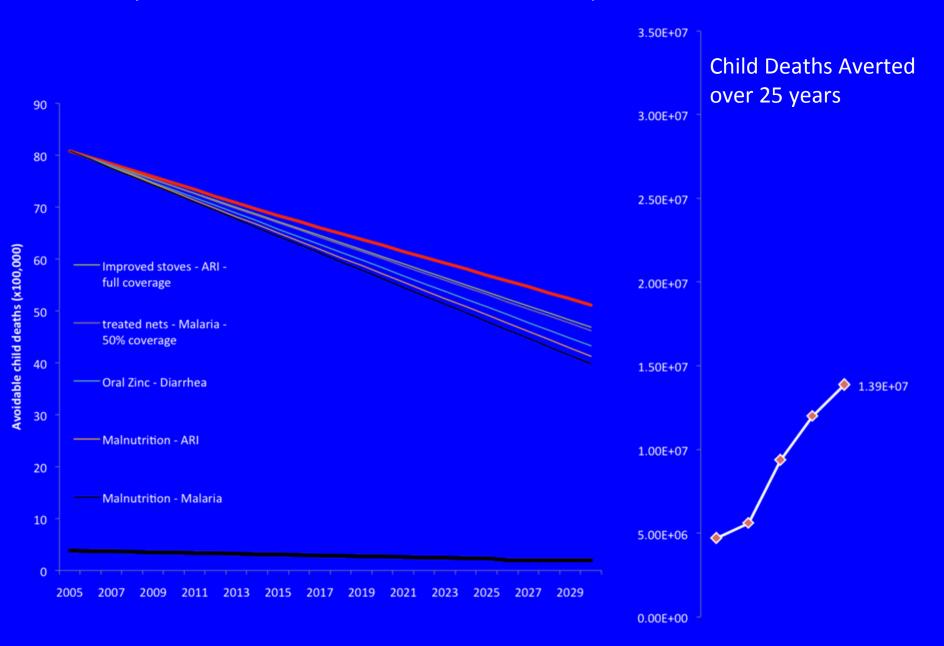


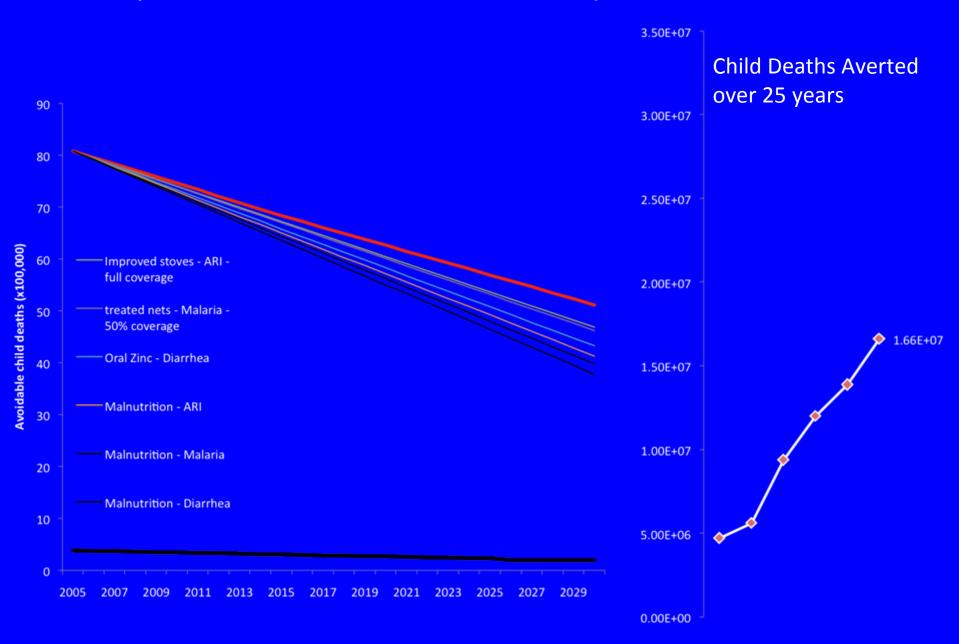
Potential Impacts of Health Interventions

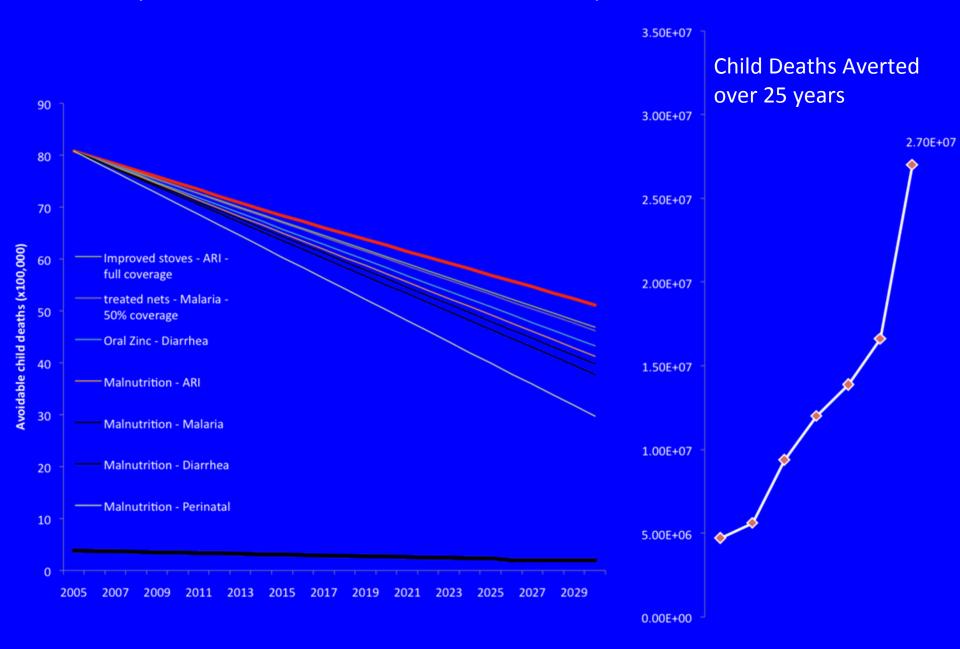


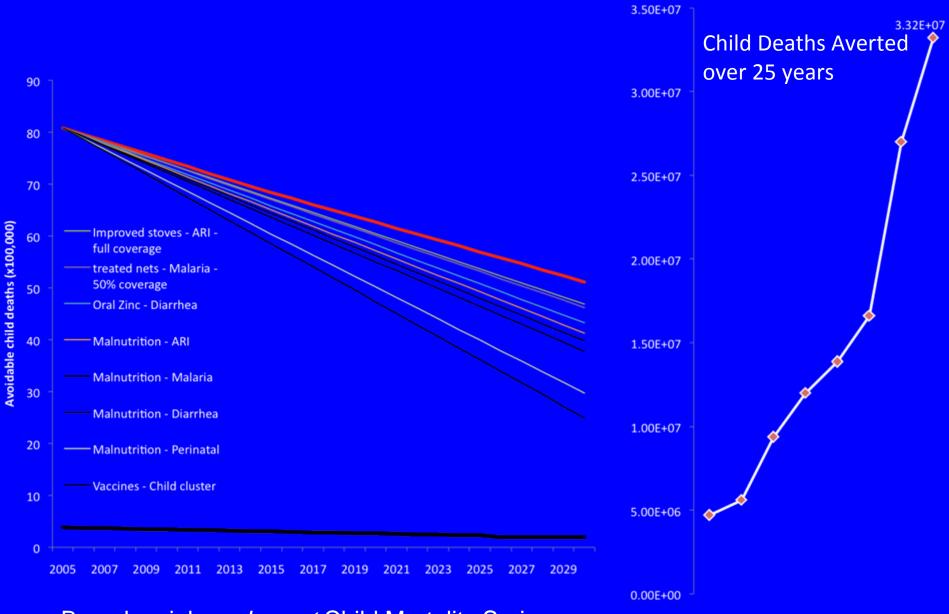




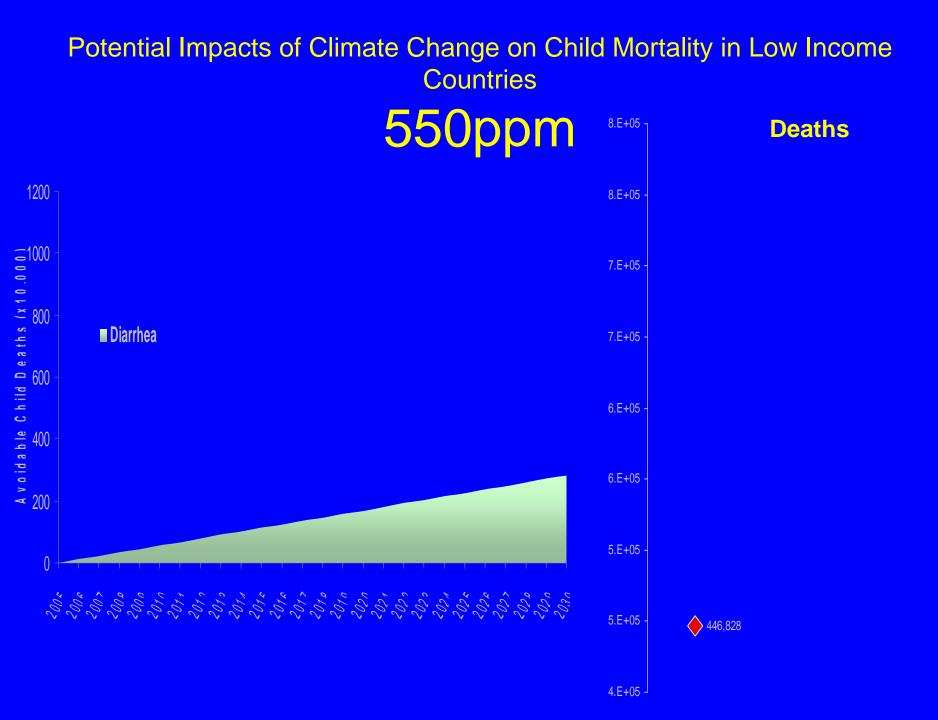


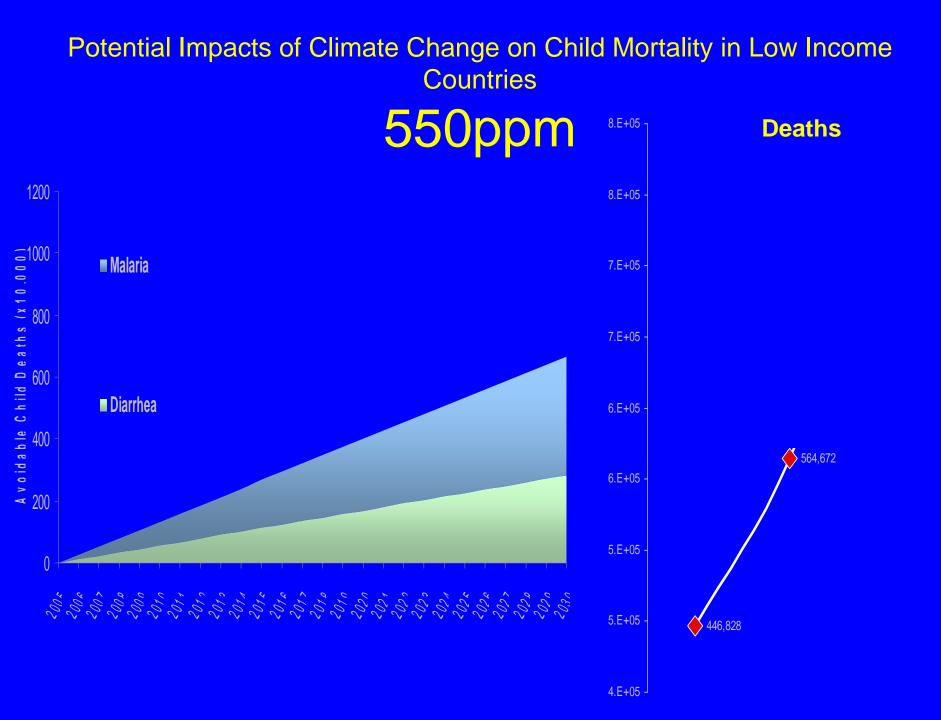


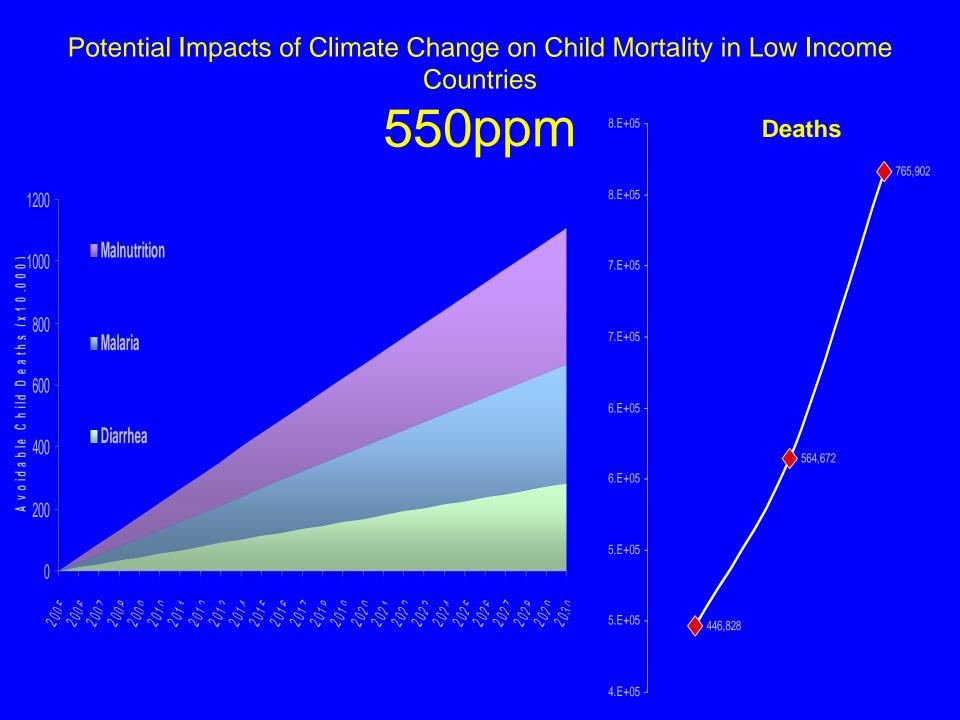




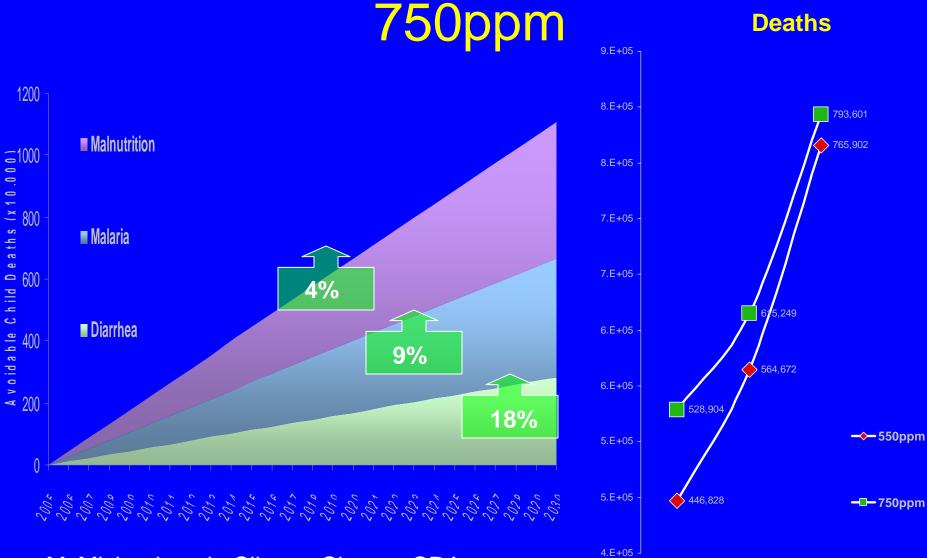
Based mainly on *Lancet* Child Mortality Series







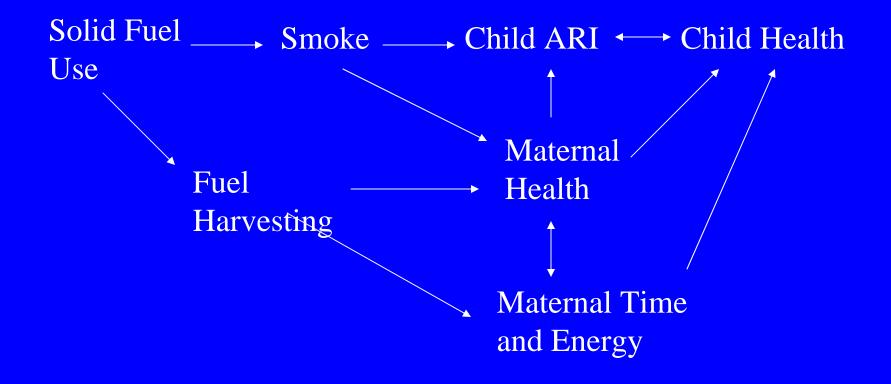
Potential Impacts of Climate Change on Child Mortality in Low Income Countries



McMichael et al., Climate Change CRA

Thank you.

The Unhealthy Mother Effect



Mark Twain on Casual Pathways

"Soap and education are not as sudden as a massacre, but they are more deadly in the long run"

- Facts Concerning the Recent Resignation, 1867