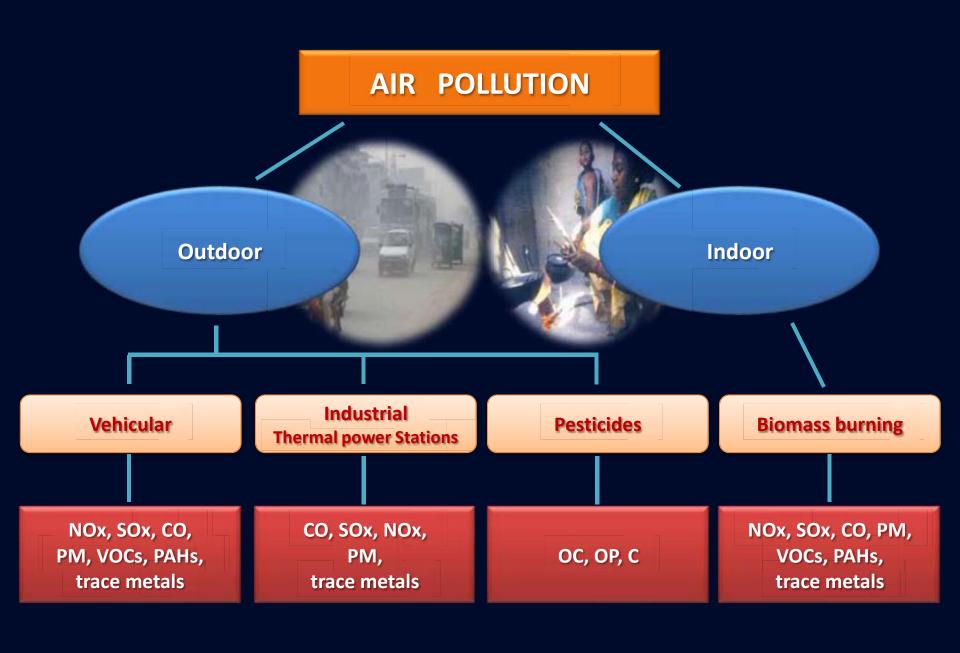
### HEALTH EFFECTS OF AIR POLLUTION

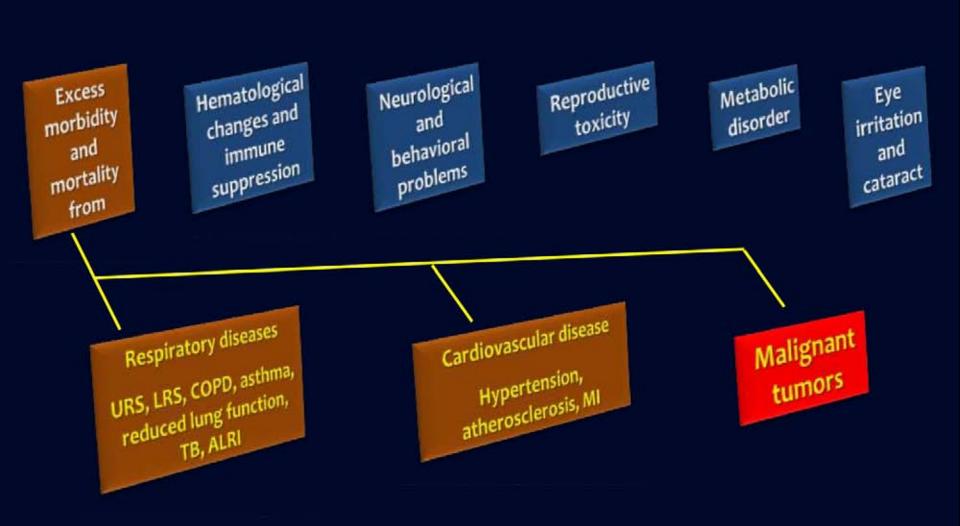
Dr. Manas Ranjan Ray

Chittaranjan National Cancer Institute Kolkata





### Health effects of air pollution



# Particulate Matter (PM) the prime concerned pollutant

PM<sub>10</sub>: diameter < 10 microns

PM<sub>2.5</sub>: diameter < 2.5 microns

Ultra Fine Particles(UFPs): diameter < 0.1 microns

Smaller the size, greater the health risk

### PM deposition in lungs and the airways

### Naso-oropharangeal region <

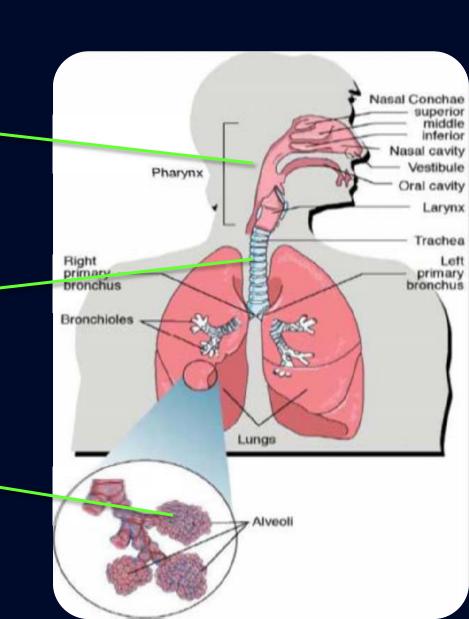
Large and water-soluble PMs are removed

### Tracheo-bronchial region 🛹

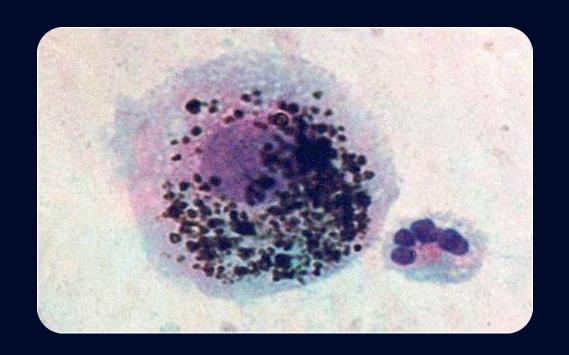
Smaller percentages of PM<sub>10</sub> and PM<sub>2.5</sub> are deposited

### Alveolar region ≼

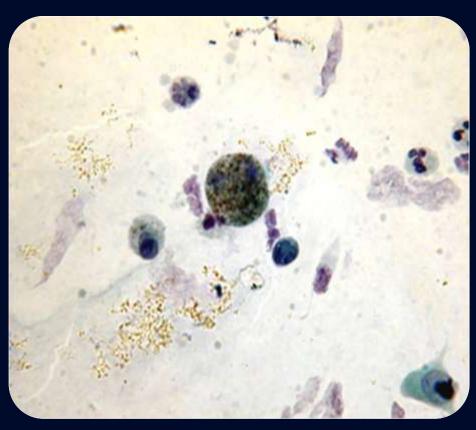
PM<sub>10</sub>, PM<sub>2.5</sub> and UFPs are deposited; a fraction of UFPs migrate to circulation



### Alveolar macrophage biomarker of pollution exposure

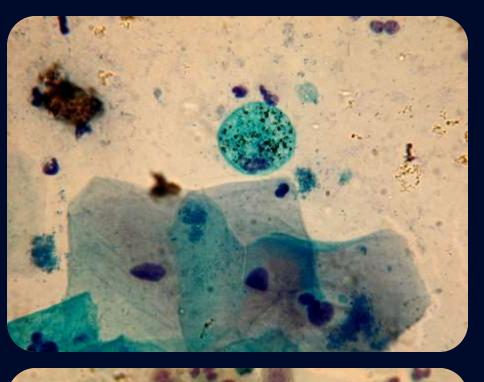


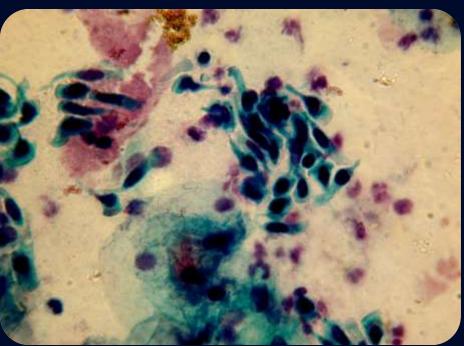
### Alveolar macrophages (AM) engulf inhaled pollutants

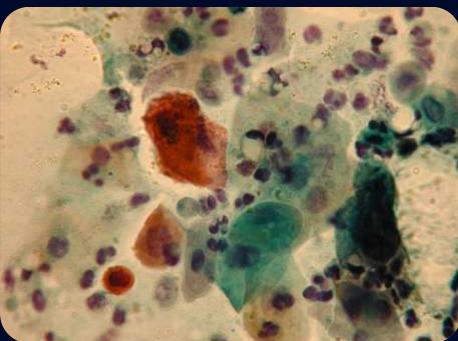


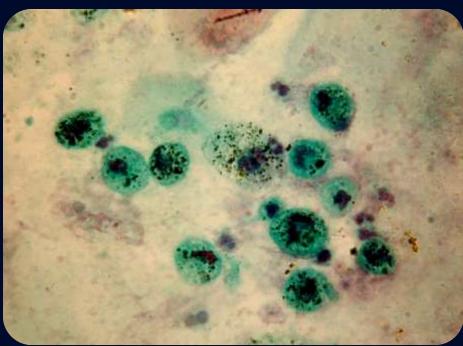


**Control Exposed** 

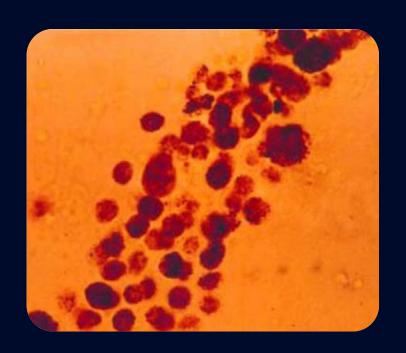








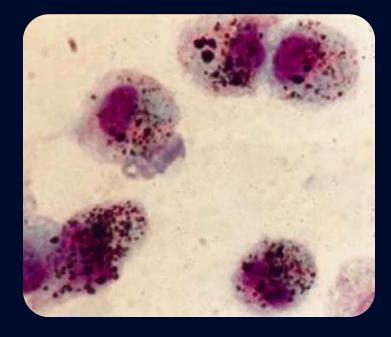
### Alveolar macrophage: the biomarker of air pollution



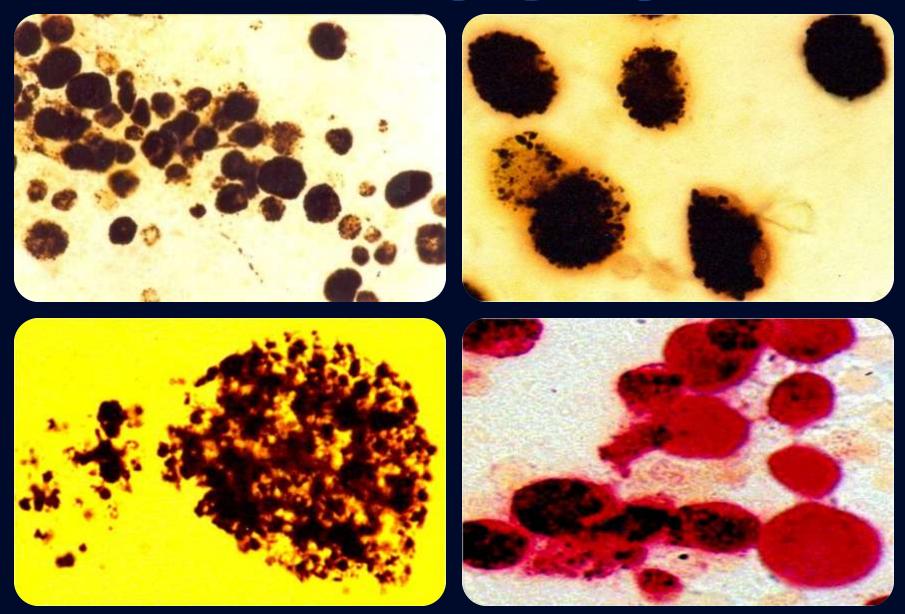
7 6 5 Delhi 5.2

Delhi 1.7
1 0 AM

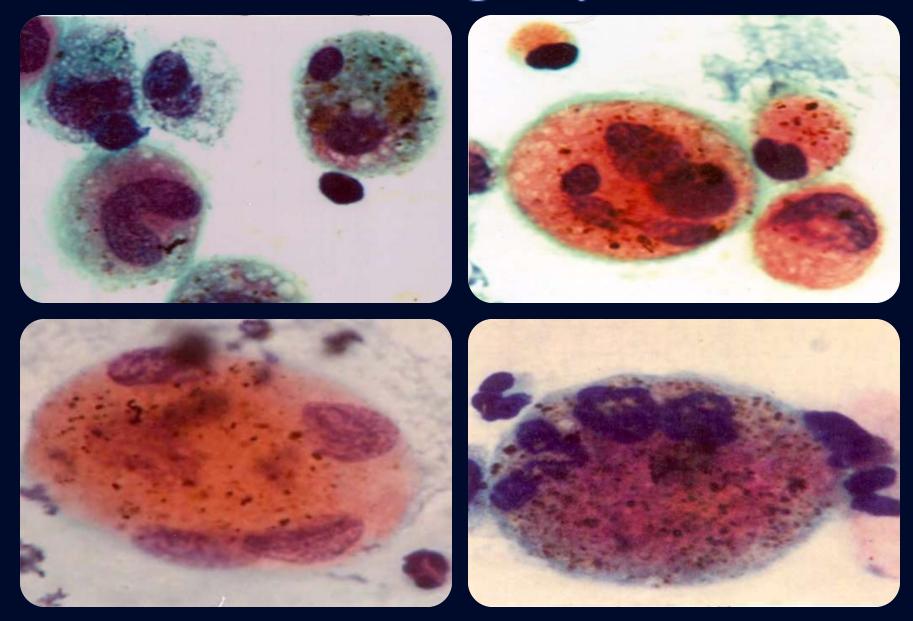
Sputum cytology of a 14-year old girl, showing abundance of particle laden AM



### Alveolar Macrophage Response



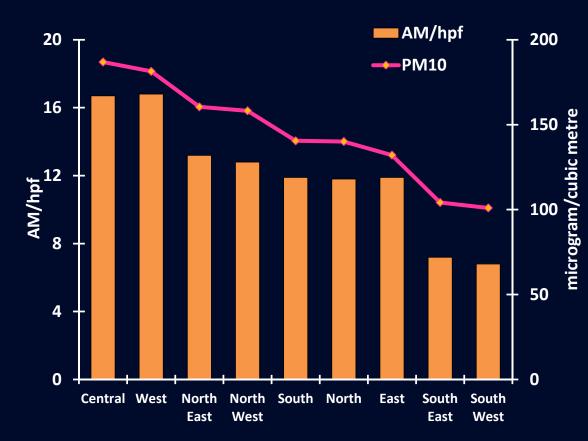
### Nuclear heterogeneity of AM



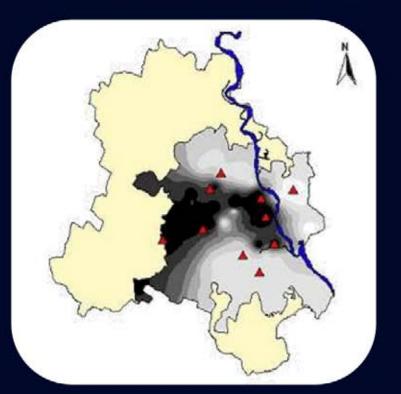
# 12 KM

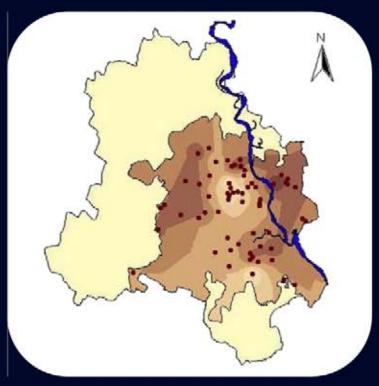
### PM<sub>10</sub> and AM count

AM count correlated positively with PM<sub>10</sub> level (rho=0.581, p<0.001)

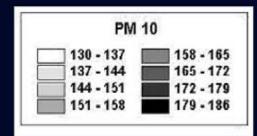


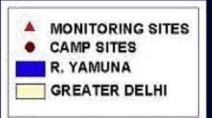
### PM<sub>10</sub> and respiratory symptoms

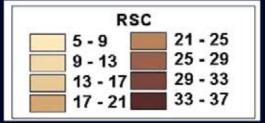




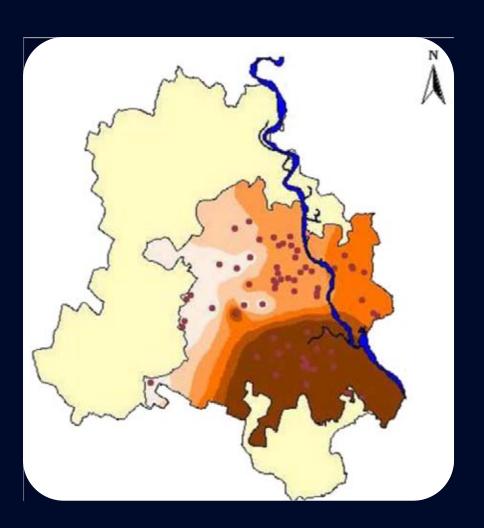


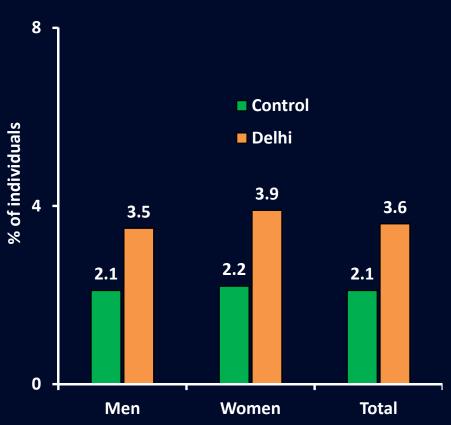




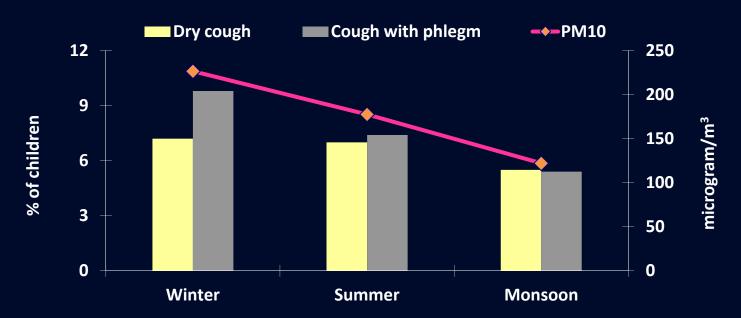


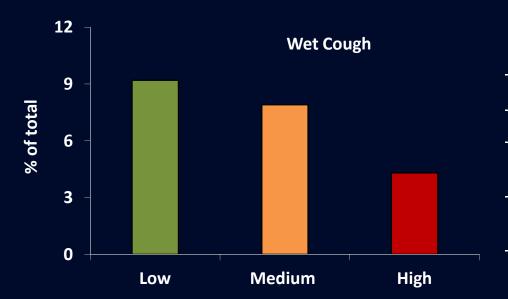
### Prevalence of bronchial asthma





### **Lower Respiratory Symptoms**





### **Inverse relation with SES**

SES	Dry cough	Wet cough	
High	1	1	
Medium	2.31* (1.96-2.73)	1.72* (1.48-2.01)	
Low	3.34* (2.83-3.95)	2.20* (1.89-2.57)	

## Strong positive association between PM<sub>10</sub> level and prevalence of LRS

PM <sub>10</sub> (μg/m³)	Dry cough	Wet cough	Wheeze	Breathless- ness	Chest discomfort	Disturbed Sleep
50-75	1	1	1	1	1	1
76-100	1.22*	1.06	0.97	1.12*	1.13*	1.26*
	(1.04-1.47)	(0.84-1.29)	(0.76-1.14)	(1.02-1.44)	(1.02-1.34)	(1.10-1.48)
101-125	1.86*	1.29*	1.04	1.34*	1.66*	1.64*
	(1.54-2.24)	(1.06-1.57)	(0.72-1.38)	(1.14-1.83)	(1.27-2.14)	(1.25-2.13)
126-150	2.20*	1.30*	1.43*	1.51*	1.94*	1.78*
	(1.81-2.68)	(1.09-1.56)	(1.12-1.77)	(1.22-1.78)	(1.53-2.44)	(1.33-2.27)
>150	3.12*	3.03*	1.67*	2.84*	2.65*	2.73*
	(2.36-3.75)	(2.53-3.62)	(1.19-2.36)	(2.31-3.47)	(2.09-3.37)	(1.89-4.32)

The results are expressed as odds ratio with 95% CI in parentheses; \*, p<0.05

### **Pulmonary function test**



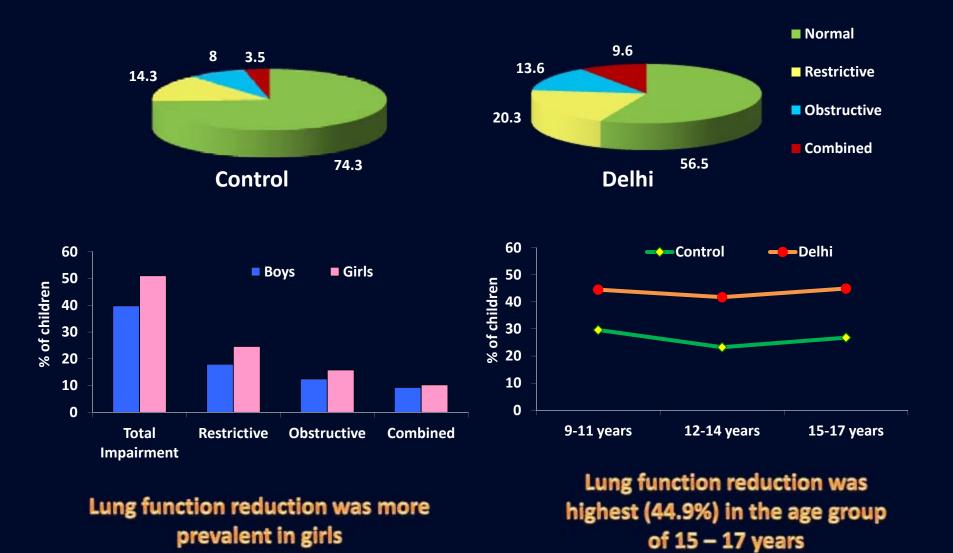
Assessed by spirometry using portable spirometer (Spirovit SP1, Schiller, Switzerland)



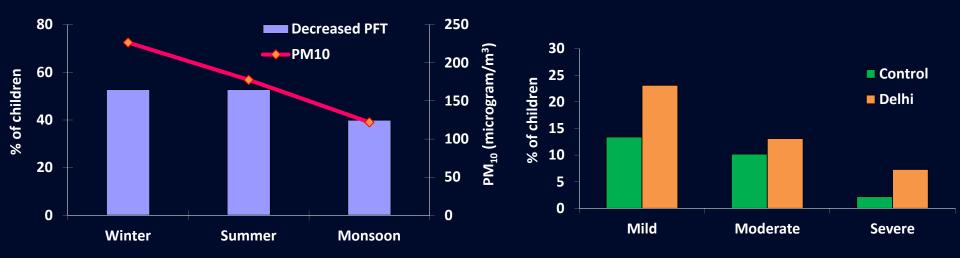
Parameters assessed : FVC, FEV<sub>1</sub>, PEFR, FEF<sub>25-75%</sub>

Lung function impairment categorized as restrictive, obstructive and combined defects & degree of impairment was ascertained

### Overall prevalence of lung function deficits



### **Reduced lung function**

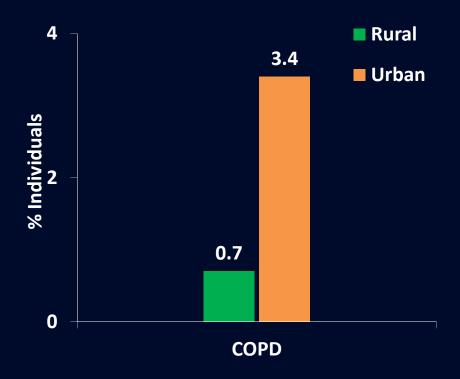




7.3% of children of Delhi had severe lung function deficits

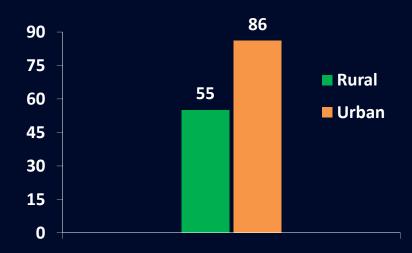
Lung function decrement was highest in North Delhi (48.5%), and lowest in South Delhi (34.3%)

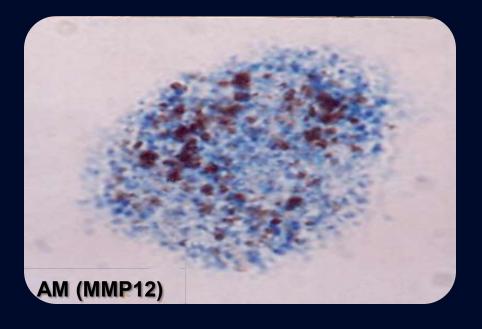
### Air pollution and COPD among never-smokers

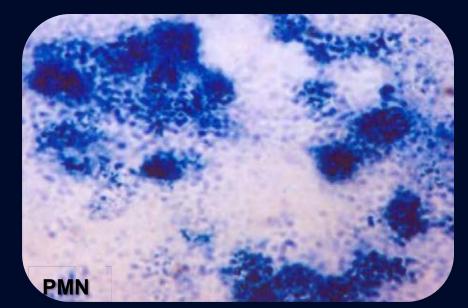


# PMN

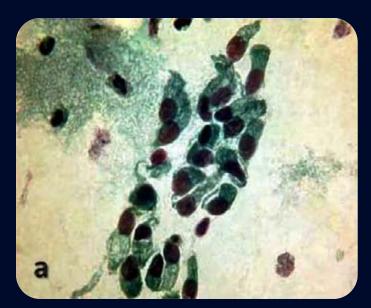
### Elastase

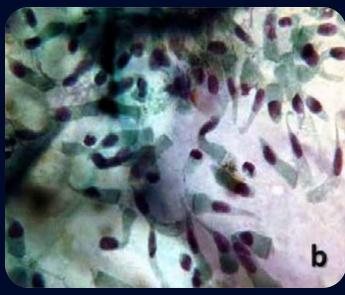






# Adverse cellular lung reactions to chronic air pollution exposures

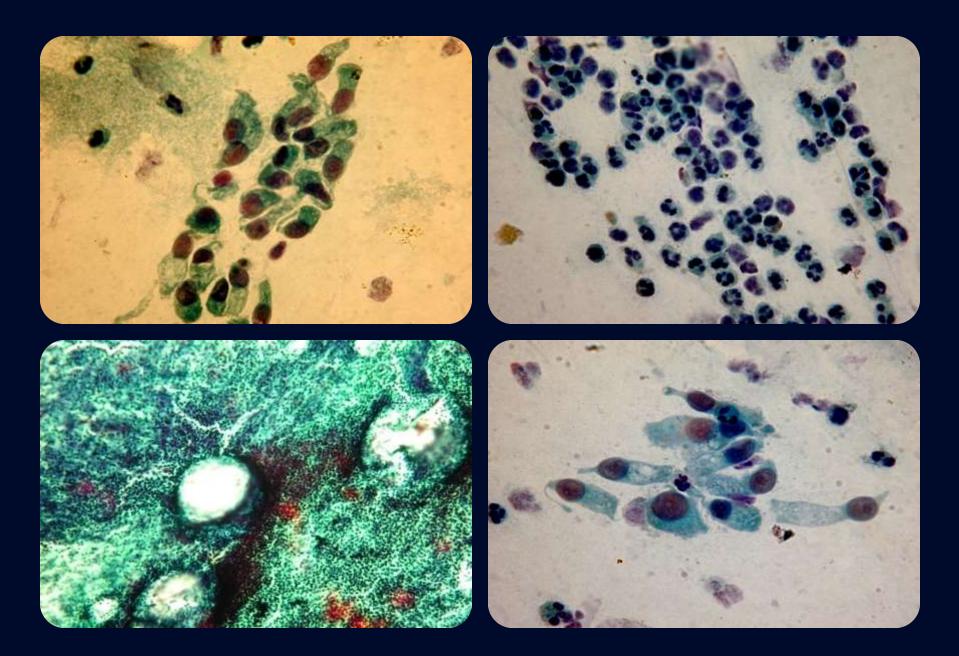




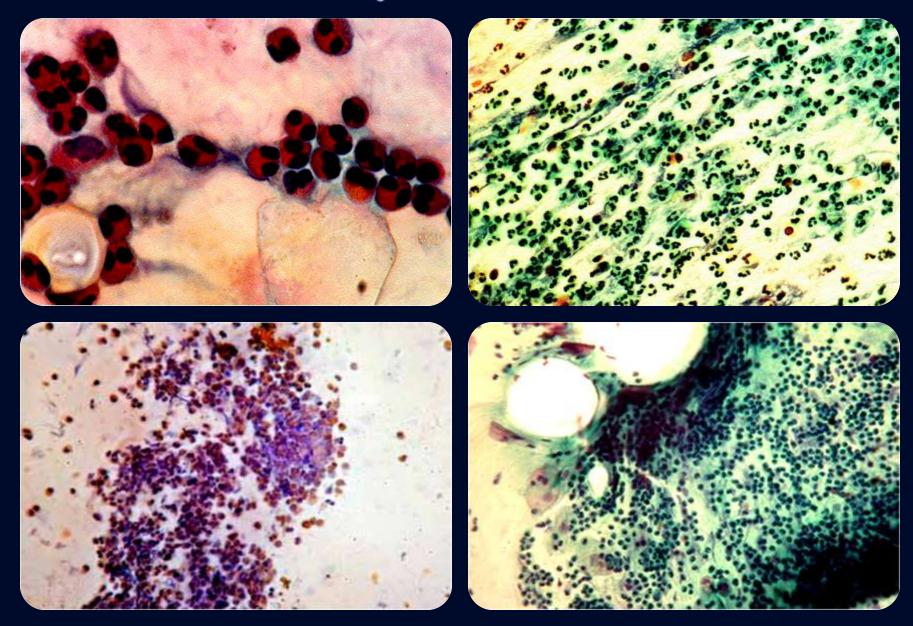




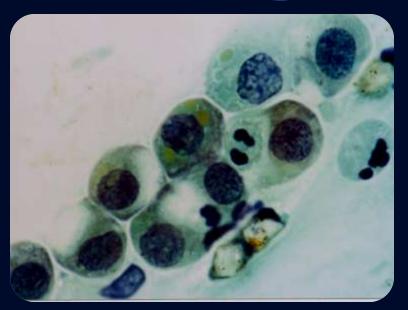




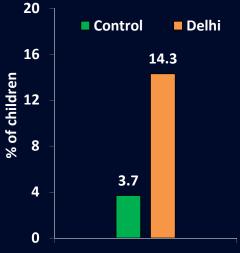
### **Airway Inflammation**

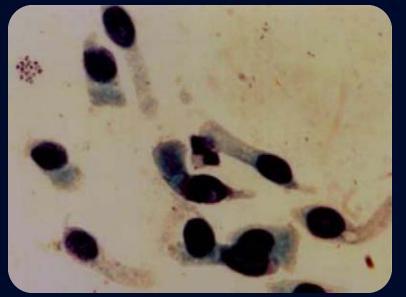


### Changes in airway epithelial cells



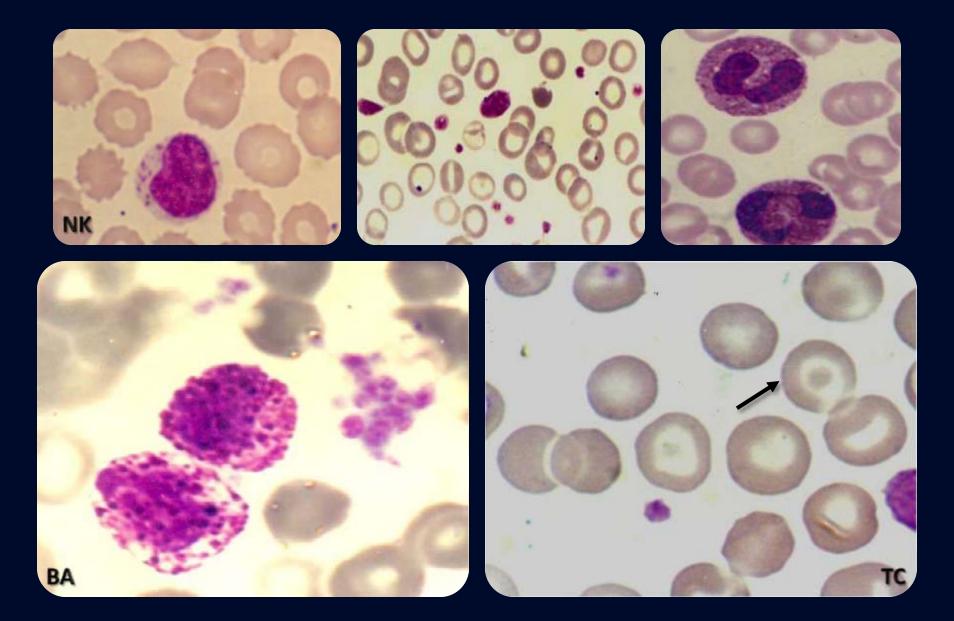
Goblet cell hyperplasia. The change indicates hypersecretion of mucous, a defence against pollutants





Aggregates of columnar epithelial cells, suggesting airway injury

### Hematological changes



### Alteration in immune status

### increased susceptibility to disease

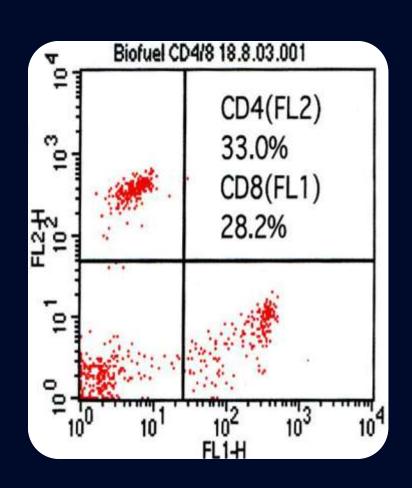
suppression of CD4+ Th cells

increase in CD 8+ Tc

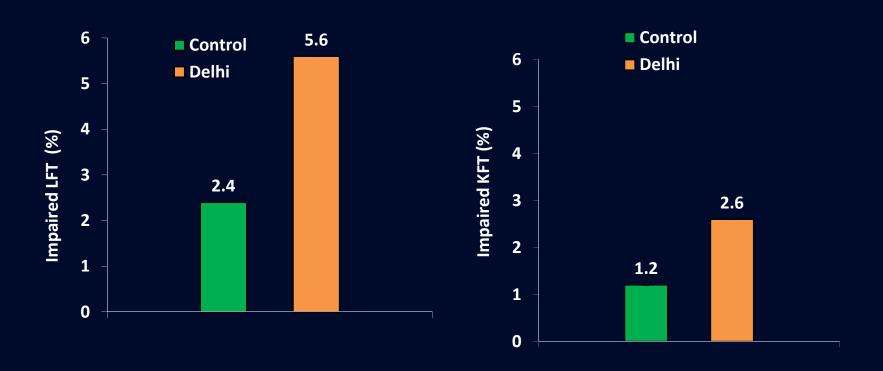
CD4:CD8 ratio 2:1 → 1:1

decrease in CD19+ B cells

increase in CD16+56+ NK cells

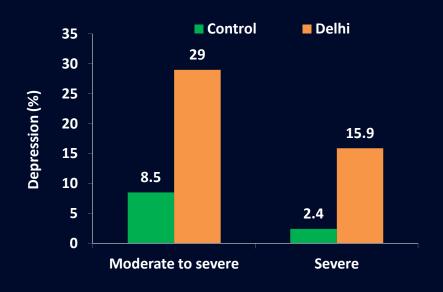


### Liver and kidney function



- 2-fold rise in liver and kidney function impairments
- 4-fold rise in diabetes: 7.4% in Delhi vs. 1.9% in control

### Neurobehavioral symptoms

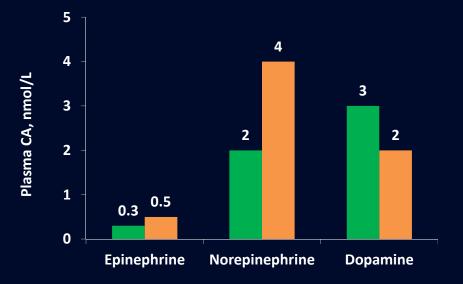


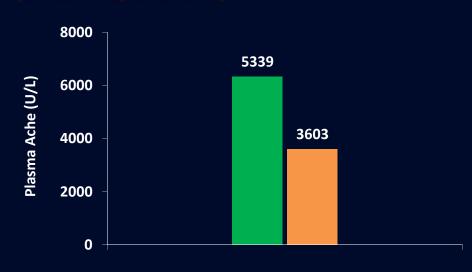


Significant alteration in plasma neurotransmitter level

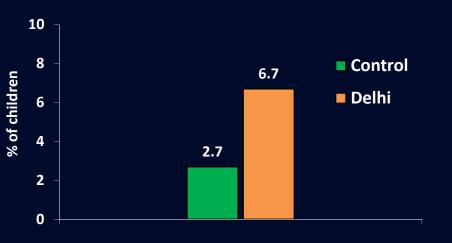
Marked fall in plasma acetylcholinesterase activity

A positive association between PM<sub>10</sub> and depression (OR=1.83)



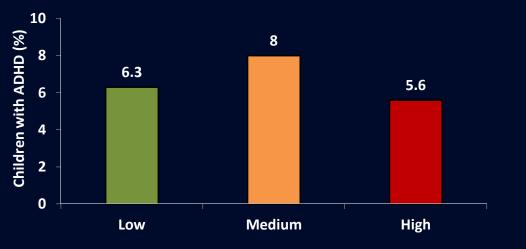


### Attention Deficit Hyperactivity Disorder (ADHD)



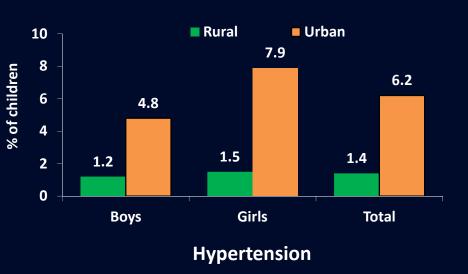


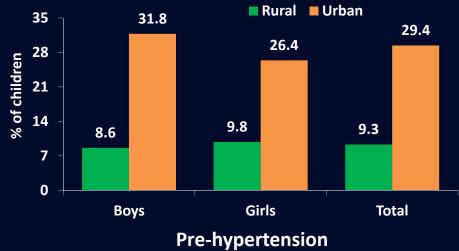
9% of the boys in Delhi had ADHD against 2% of the girls, giving a male: female ratio of 4.5:1

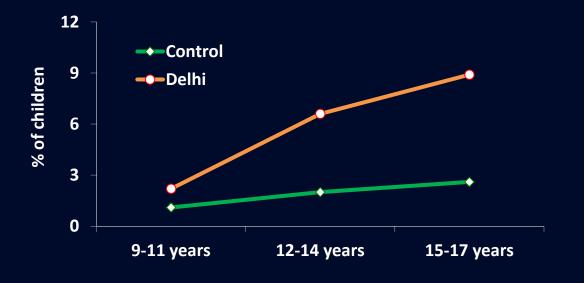


Children with ADHD are impulsive, forgetful, restless, prone to fail, unable to follow tasks, unpredictable and moody

### **Hypertension - I**

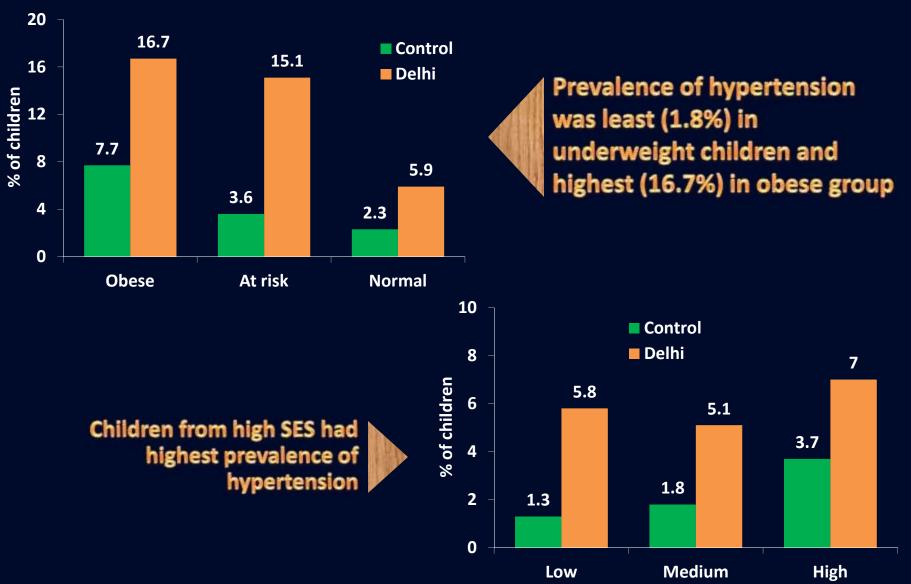




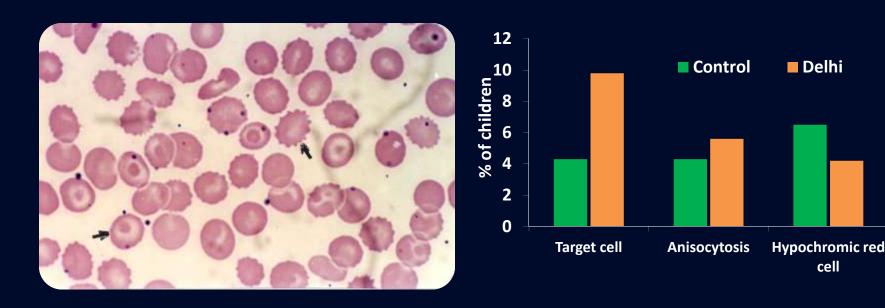


Prevalence of hypertension increased progressively with age

### **Hypertension - II**



### Hematological alterations- I

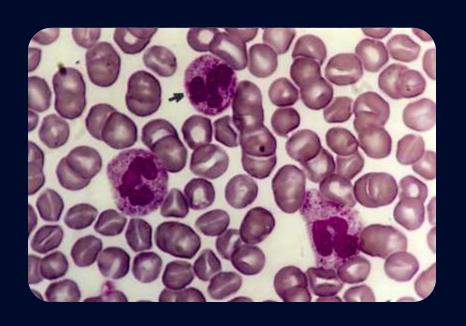


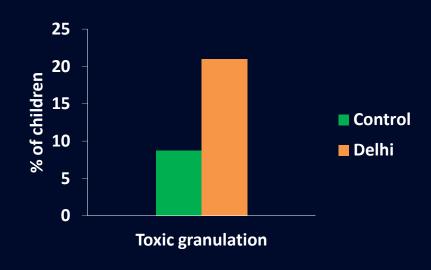
9.8% of Delhi's children had abundance of 'target' cells in their peripheral blood compared with 4.3% of controls (p<0.001)

cell

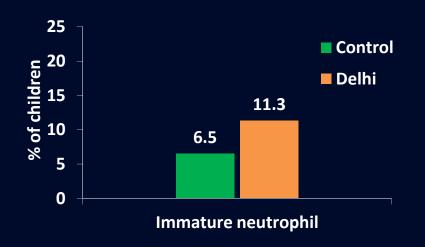
Their presence in circulation in excess signifies liver problem

### Hematological alterations- II

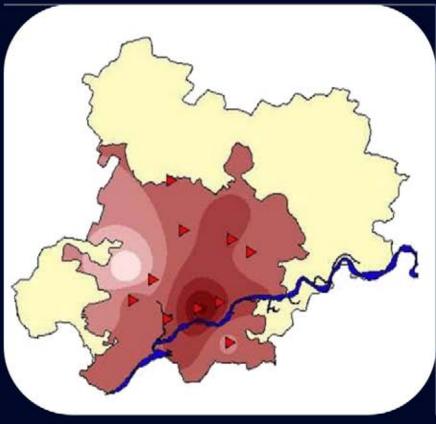


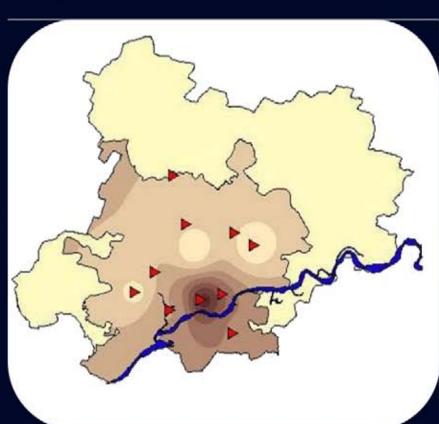


Higher prevalence of circulating immature neutrophil and toxic granulation in neutrophil in Delhi's school children suggests greater risk of infection and inflammation



# Benzene and B[a]P distribution in Delhi





Benzene

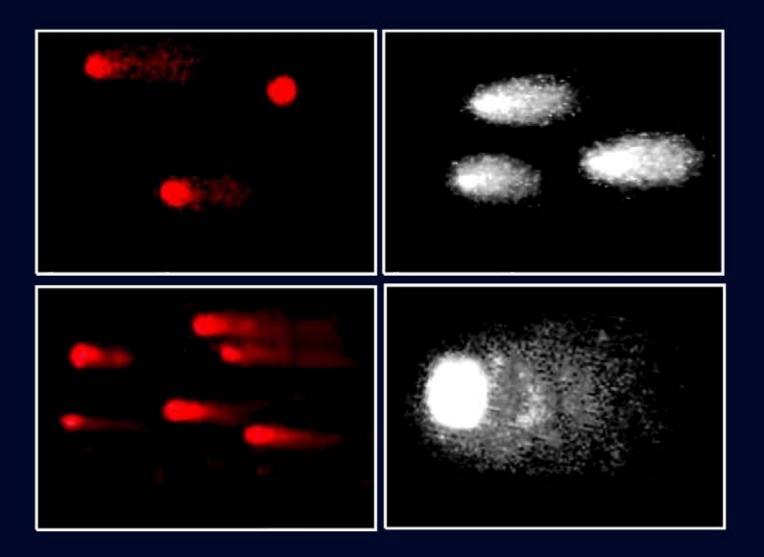
Mean concentrations of

B[a]p

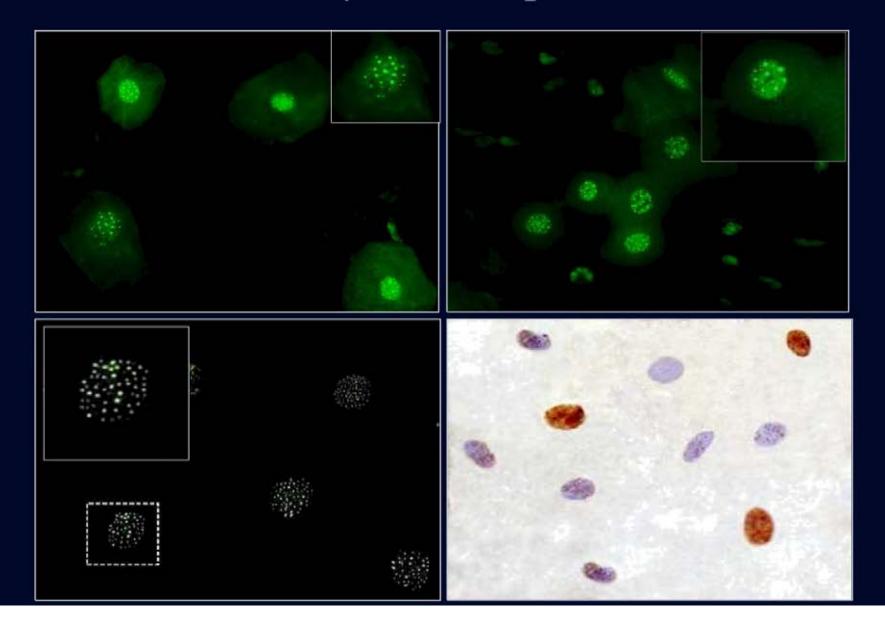
Benzene: 21.4 μg/m³

Benzo[a]pyrene: 3.82 ng/m³

#### Metabolically active carcinogens induce DNA damage

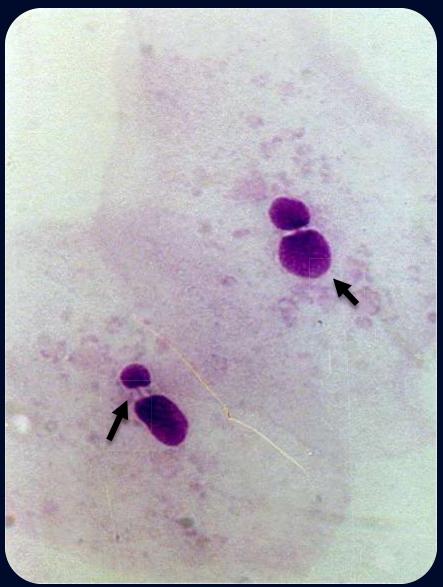


# DSB: γH2AX expression



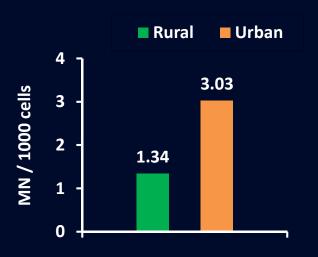
## MN (Chromosomal breaks)



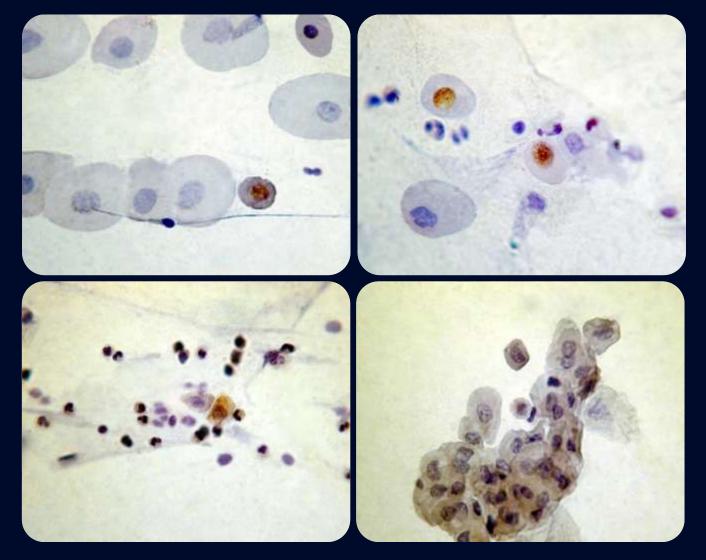


### Chromosomal damage





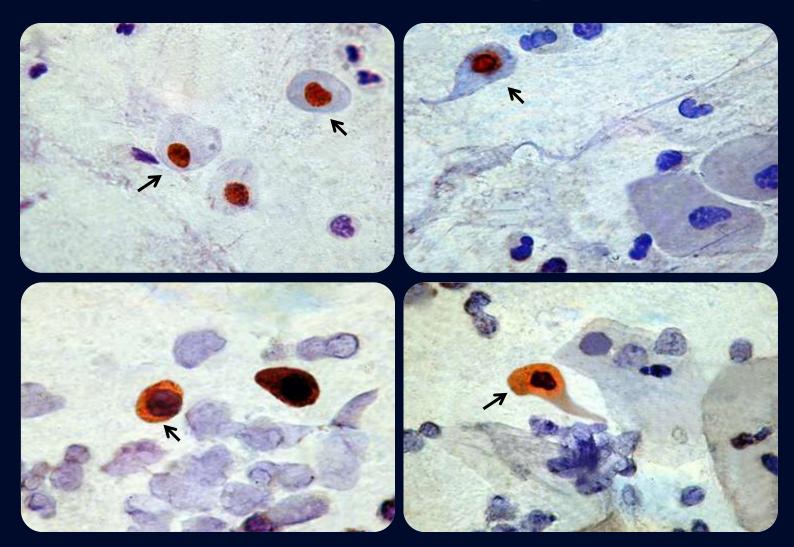
#### Deficiency in DNA repair



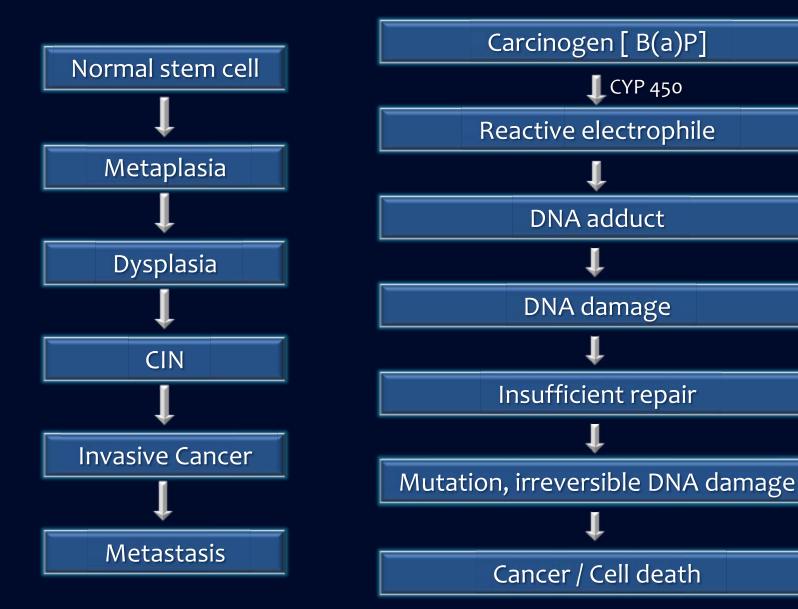
NHEJ, MMR, BER in airway epithelial cells

# **Up-regulation of Akt**

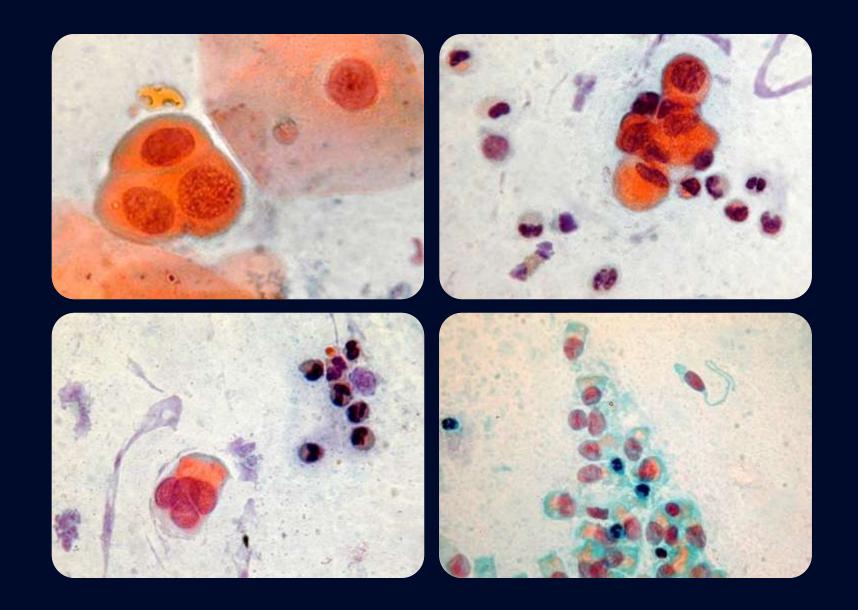
**Proliferation advantage** 



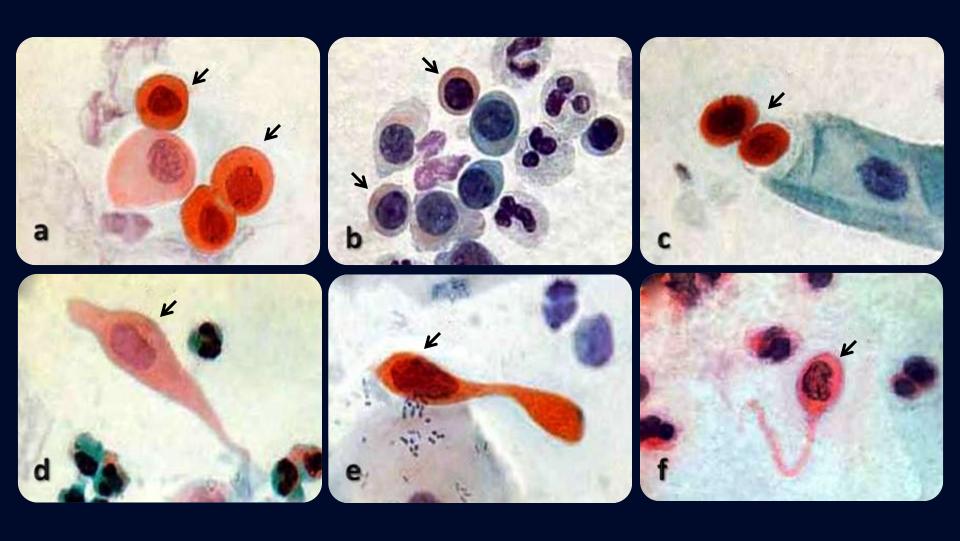
#### Journey towards cancer

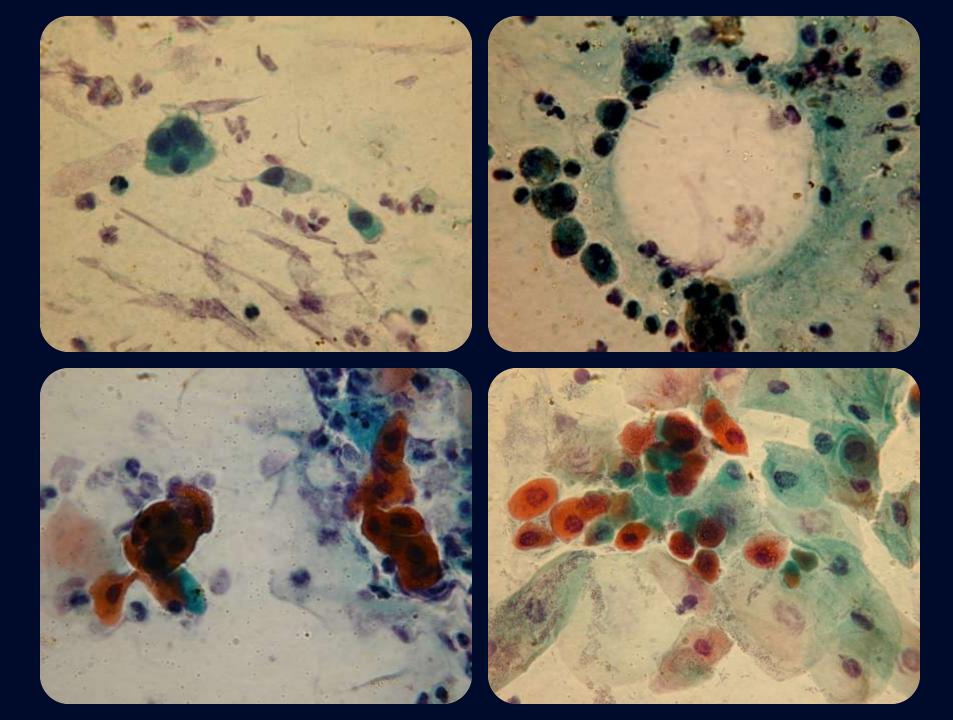


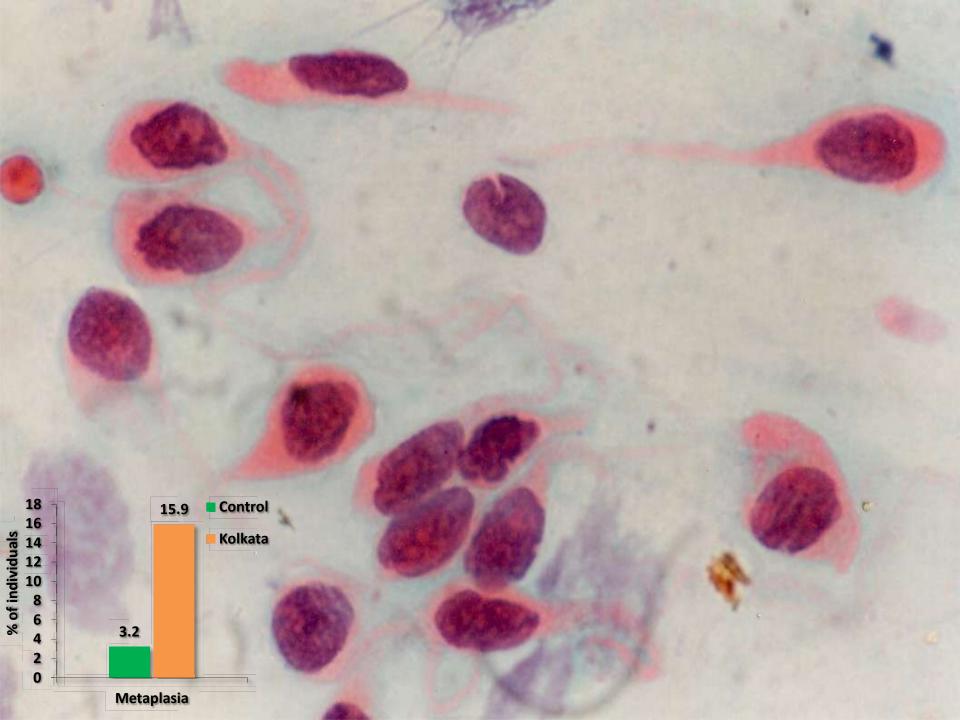
#### Airway cells following DNA damage, precancerous changes



# Metaplasia and dysplasia of airway cells among never-smoking biomass-using women









#### Cancer incidence in rural and urban Delhi

Site of cancer	Rural	Urban
Male (overall)/100,000	55.2	116.9
Lung (%)	6.5	13.8
Oral (%)	8.0	11.4
Larynx (%)	4.0	7.9
Female (overall)/ 100,000	47.7	116.7
Breast (%)	7.8	30.2
Uterus (%)	10.3	17.5
Ovary (%)	3.3	8.5
Gall bladder (%)	3.5	7.4

#### The Study Team



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Dr. M. Banerjee

Dr. S. Siddique

Dr. S. Chakraborty

Dr. A. Dutta

Dr. N. K. Mondal

Dr. A. Banerjee

Dr. D. Das

Mrs. S. Choudhury

Mr. P. B. Paul

# THANK YOU