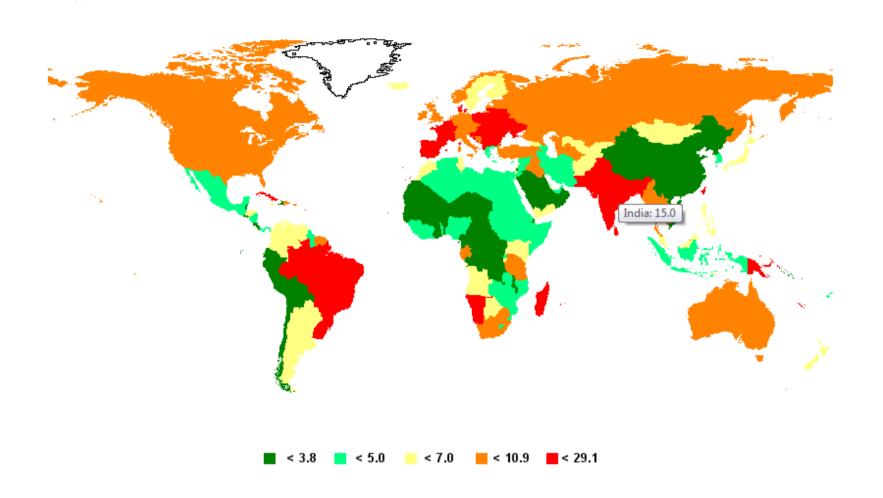
# Head & Neck Cancer: The Current Scenario and Epidemiology

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Estimated age-standardised incidence rate per 100,000 Lip, oral cavity, Larynx, Other pharynx: both sexes, all ages



# 6.4 lakh H & N Ca diag worldwide/yr 1.5 lakh India/year ~20 %



## About Cancer Registry in India

- The National Cancer Registry Programme (NCRP) was commenced by the Indian Council of Medical Research (ICMR) with a network of cancer registries across the country in December 1981.
- Population based cancer registries record all the new cancer cases occurring in a defined geographic area, with epidemiological and public health aspects in mind.

Source: www.ncrpindia.org

 The hospital based cancer registries record information on cancer patients attending a particular hospital, with focus on clinical care and hospital administration.

 At present, there are 23 PBCRs and 7 HBCRs under the NCRP network.



Table 9.1: Number (#) and Proportion (%) of Head and Neck Cancers
Relative to All Sites of Cancer (2007-2011)

Registry	All Cites	Males	0/	All Cites	Females	0/
	All Sites	#	%	All Sites	#	%
Mumbai	22580	6805	30.1	18528	1673	9.0
Bangalore	11273	3532	31.3	13125	1822	13.9
Chennai	15731	4427	28.1	17499	1832	10.5
Thi'puram	19219	4798	25.0	18809	1726	9.2
Dibrugarh	2895	1211	41.8	2276	329	14.5
Guwahati	6803	2830	41.6	4679	702	15.0
Chandigarh	2643	598	22.6	2092	100	4.8
Total	81144	24201	29.8	77008	8184	10.6

Report from Hospital Based Cancer Registries
 Source: ncrpindia.org



Table 9.3: Number (#) and Relative Proportion (%) of Specific Head and Neck Sites
Relative to All Head and Neck Cancers (2007-2011)

Sites of Cancer	Mun	nbai	Bang	alore	Che	nnai	Thi'p	uram	Dibr	ugarh	Guw	ahati	Chan	digarh
onde of dancer	#	%	#	%	#	%	#	%	#	%	#	%	#	%
MALES														
Tongue	1603	23.6	754	21.3	1061	24.0	1214	25.3	164	13.5	472	16.7	192	32.1
Mouth	2776	40.8	731	20.7	1351	30.5	1593	33.2	257	21.2	545	19.3	94	15.7
Nasopharynx	172	2.5	61	1.7	155	3.5	127	2.6	31	2.6	75	2.7	27	4.5
Other Pharynx	1413	20.8	1539	43.6	1261	28.5	1098	22.9	653	53.9	1356	47.9	145	24.2
Larynx	841	12.4	447	12.7	599	13.5	766	16.0	106	8.8	382	13.5	140	23.4
Head and Neck	6805	100.0	3532	100.0	4427	100.0	4798	100.0	1211	100.0	2830	100.0	598	100.0



Fig. 9.2: Stack (100%) Diagram Showing Proportion (%) of Specific Head and Neck Cancer Sites
Relative to All Head and Neck Cancers (2007-2011)

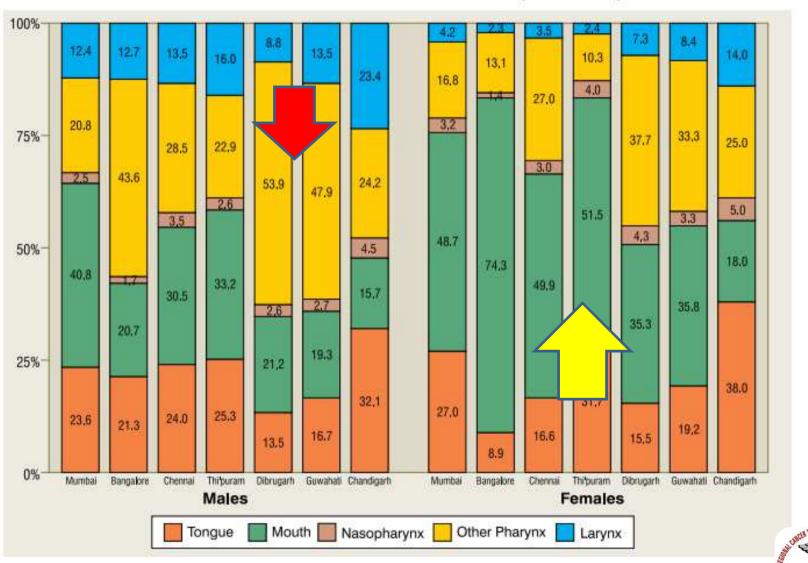
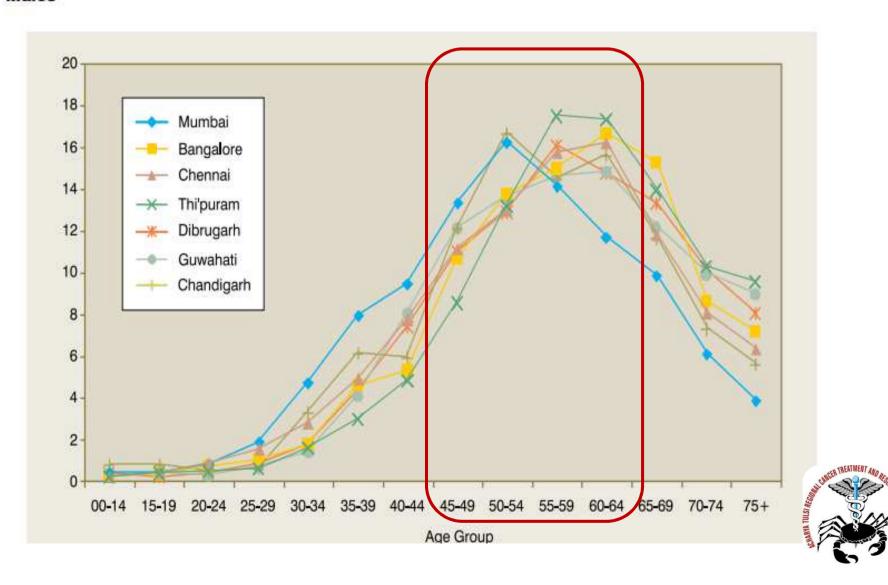


Fig. 9.3: Relative Proportion (%) of Head and Neck Cancers by Five-Year Age Group (2007-2011)

#### Males



### Cancer Burden ahead

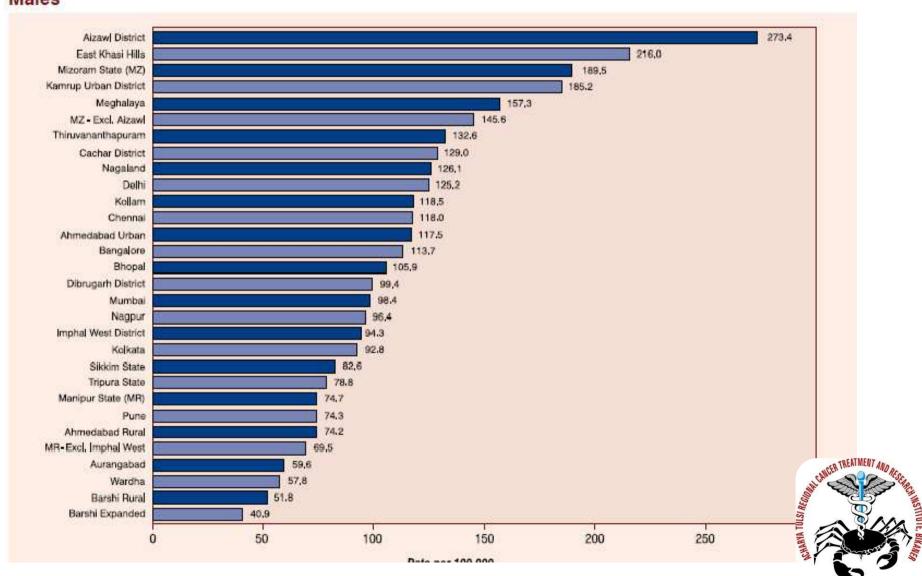
Table 5.1(b): Projected Cases at India level for Selected Sites and Selected Time Periods (2013, 2014, 2015 and 2020)

#### **Both Sexes**

	w.	1,26,304	ii i	i i	1,68,637
ICD-10	Site Name	2013	2014	2015	2020
C00-C96	All Sites	1086783	1117269	1148692	1320928
C01-02	Tongue	44449	46614	48888	62099
C03-06	Mouth	63627	66836	70228	90342
C12-13	Hypopharynx	16238	16224	16213	16196
C15	Oesophagus	37909	37623	37344	36058
C16	Stomach	35531	36176	36837	40419
C18	Colon	28883	30312	31816	40601
C19-20	Rectum	27960	29173	30442	37720
C22	Liver	26738	27781	28866	35003 35003

Fig. 7.1: Comparison of Age Adjusted Incidence Rates (AARs) of all PBCRs
ALL SITES (ICD-10: C00-C97)

#### Males



#### TOP TEN CANCERS: MALE

			MALE			
Bangalore	Barshi	Bhopal	Chennai	Delhi	Mumbai	Ahmedabad
Stomach(9.3)	Hypopharynx(3. 8)	Lung(10.1)	Lung(12.4)	Lung(13.5)	Lung(8.7)	Mouth(6.9)
Lung(8.5)	Esophagus(2.6)	Mouth(8.8)	Stomach(11.9)	Prostate(10.4)	Mouth(6.8)	Tongue(7.0)
Esophagus(7.0)	Larynx(2.8)	Tongue(8.9)	Esophagus(7.9)	Larynx(8.5)	Prostate(6.8)	Lung(6.6)
Prostate(6.8)	Mouth(3.4)	Esophagus(5.4)	Mouth(6.3)	Tongue(6.6)	Larynx(5.5)	Hypopharynx(5. 3)
Brain(3.9)	Stomach(2.6)	Larynx(4.7)	Tongue(5.7)	NHL(5.4)	Esophagus(5.5)	Esophagus(3.3)
NHL(4.1)	Prostate(1.6)	Prostate(4.7)	Larynx(4.9)	Bladder(6.5)	Tongue(4.8)	Larynx(2.7)
Liver(4.4)	Penis(2.0)	Hypopharynx(4. 1)	Prostate(5.0)	Mouth(5.5)	NHL(4.4)	Myel Leuk(1.8)
Hypopharynx(3	Oth Skin(1.6)	NHL(3.2)	NHL(4.1)	Brain(4.2)	Stomach(4.9)	Tonsil(1.9)
Larynx (3.8)	Tongue(1.9)	Liver (2.5)	Hypopharynx(4. 5)	Esophagus(5.1)	Liver(4.6)	Brain(1.5)
Colon (3.2)	NHL(1.9)	Myel Leuk(1.9)	Rectum(3.7)	Stomach(3.5)	Brain(3.1)	Prostate(1.7)

#### TOP TEN CANCERS: FEMALE

			FEMALE			
Bangalore	Barshi	Bhopal	Chennai	Delhi	Mumbai	Ahmedabad
Breast(30.9)	Cervix(22.8)	Breast(24.6)	Breast(33.0)	Breast(31.4)	Breast(29.3)	Breast(9.2)
Cervix (18.8)	Breast(9.4)	Cervix(17.7)	Cervix (22.3)	Cervix (17.4)	Cervix(13.4)	Cervix (7.9)
Ovary(6.2)	Esophagus(4.0)	Ovary(6.5)	Stomach(5.6)	Ovary(8.3)	Ovary(6.5)	Ovary(3.4)
Esophagus(6.5)	Ovary(3.8)	Gall bladder(4.3)	Ovary(5.4)	Gall bladder(7.4)	Esophagus(3.7)	Esophagus(1.7)
Mouth(5.9)	Lung(2.5)	Esophagus(4.2)	Esophagus(5.0)	Corpus uteri(4.4)	Mouth(3.5)	Mouth(1.6)
Stomach(4.9)	Rectum(0.7)	Mouth(3.7)	Mouth(4.7)	Lung(3.5)	Lung(3.0)	Myeloid Leukemia(1.0)
Corpus uteri(4.3)	Oth Skin(1.1)	Brain(1.9)	Thyroid(2.9)	NHL(2.9)	NHL(2.7)	Rectum(0.9)
NHL(3.5)	Mye Leuk(0.9)	Myeloid Leukemia(2.2)	Lung(3.0)	Brain (2.6)	Corpus uteri(2.8)	Vulva(0.8)
Thyroid(2.9)	Corpus Uteri(0.9)	Corpus Uteri(2.3)	Brain(2.2)	Thyroid(2.6)	Gall bladder(2.6)	Bone(0.7)
Lung(3.0)	Liver(0.8)	Lung(2.1)	Corpus Uteri(2.3)	Esophagus(3.1)	Stomach(2.4)	Brain ( TREATMENT AND ASSESSMENT ASSESSMENT AND ASSESSMENT AND ASSESSMENT AND ASSESSMENT AND ASSESSMENT AND ASSESSMENT ASSESSMENT ASSESSMENT ASSESSMENT AND ASSESSMENT ASSESSMENT ASSESSMENT ASSESSMENT ASSESSMENT ASSESSMENT AND ASSESSMENT ASSES

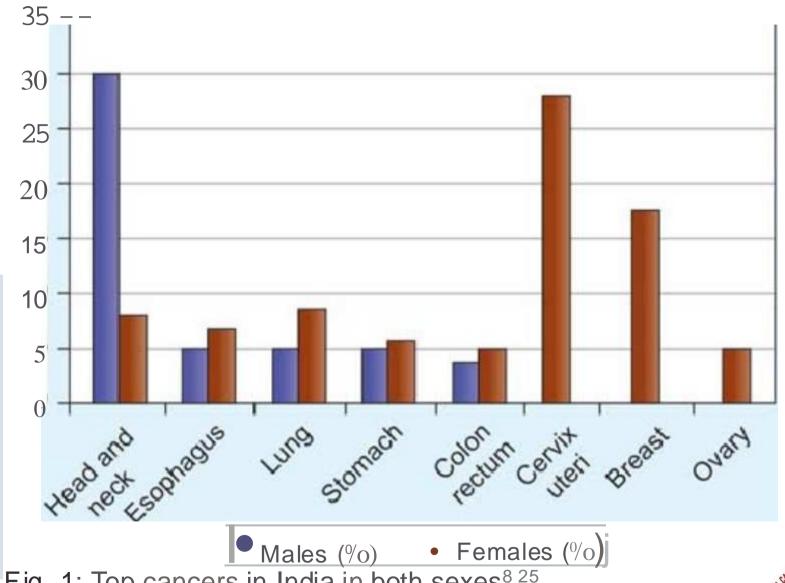


Fig. 1: Top cancers in India in both sexes<sup>8 25</sup>

#### **H&N** Cancer: Presentation

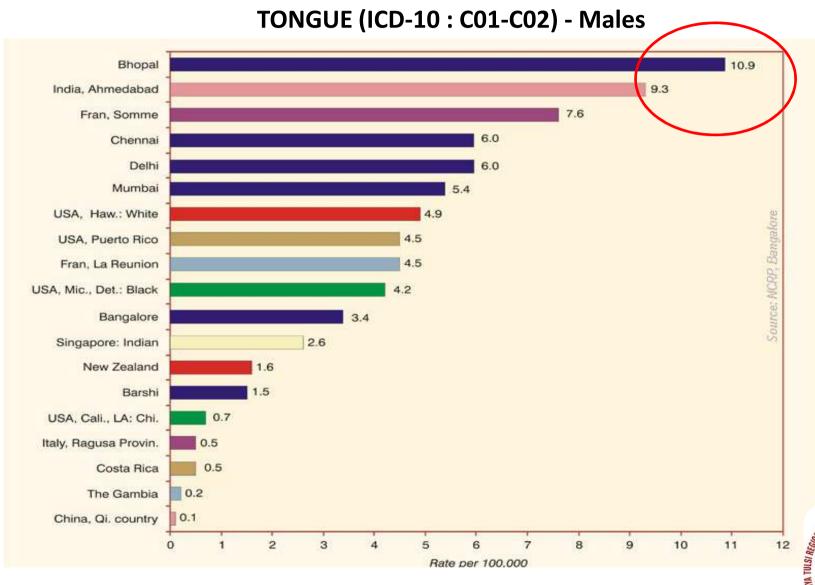
Stage <sup>1</sup>	Rate(%)	TNM <sup>2</sup>	Rate(%)	
At Presentation				
Early (III)	1530	T12	20	
LA (IIIIV)	6080	T34	80	
Metastatic	215	N01	54	
Posttherapy				
Dist Mets (+SMN)	1530	N23	46	

- 1. Choong N. CA 2008;58:32-53.
- 2. AIIMS data. JLO 2007; 121:49-56.

Usual age of diagnosis: after 40 years, except for salivary gland and nasopharyngeal cancers (younger age).



## International Comparisons of Age Adjusted Incidence Rates with that of PBCRs under NCRP



Source: Cancer Atlas (www.canceratlasindia.org)

#### **Risk Factors**

- Tobacco use
- Alcohol
- The effects of alcohol and tobacco may be synergistic.
- HNC patients: increased risk for second primary tumor (SPT), both within the head and neck and elsewhere (e.g., esophageal and lung cancers), known as field cancerisation.
- Occupational exposures to wood and leather dust: Sinonasal tract tumors.

#### Clinical Presentation

- Persistent hoarseness
- Palpable mass in neck
  - Branchial cleft cysts rarely present later than young adulthood
  - Neck mass in persons older than 40 yrs of age should be considered a malignancy until proven otherwise
- Ear infection or pain
- Altered oral sensations or persistent sore throat
- Lesions in mouth
  - Erythroplasia (early red lesions)
  - Leukoplakia (white lesions)
  - Persistent mass or ulcer (usually oral cavity)
- Difficulties in chewing, swallowing, or moving the tongue or jaws

Chin D, et al. Expert Rev Anticancer Ther. 2006;6:1111-1118. NCCN Clinical Practice Guidelines. Head and Neck Cancers. V2.2011. ACS. Oral Cavity and Oropharyngeal Cancer, Laryngeal and Hypopharyngeal Cancer.

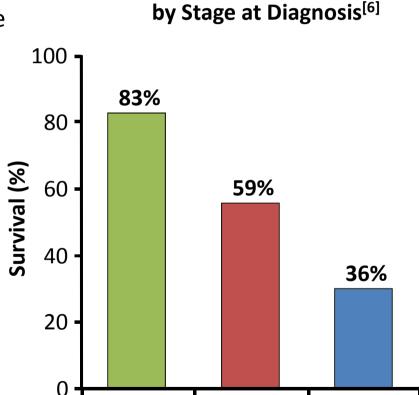
#### **Evaluation**

- Inspection and palpation
- Biopsy of any suspicious mucosal surface
- Imaging
  - CT, MRI
  - PET/CT of value in identifying neck disease and unknown primaries
  - CT of chest if there are neck nodes and no PET/CT as lung metastases common first distant site
  - New cystic lesion in the neck unlikely to be recent onset branchial cleft cyst in an adult
- FNA of lymph node
- Examination under anesthesia
  - Full evaluation of the areas at risk

NCCN Clinical Practice Guidelines. Head and Neck Cancers. V2.2011. ACS. Oral Cavity and Oropharyngeal Cancer, Laryngeal and Hypopharyngeal Cancer.

# HNSCC: Survival Rates by Stage of Disease

- High cures rates are achieved for localized and loco-regional disease using:
  - Surgery
  - Radiation
  - Chemoradiation
  - ± Induction chemotherapy
- Survival rates for recurrent/ metastatic disease remain very poor
- Better treatment options are necessary



Localized

Regional

**Distant** 

5-Yr Relative Survival Rate

Level of Evidence for Smoking-Attributable Cancers
According to the United States Office of the
Surgeon General by Cancer Site and Yearly SmokingAttributable Mortality at Sites with Available
Estimates, United States, 2004

	Cancer Site	Yearly Smoking- Attributable Mortality
Evidence Sufficient to Infer Causal Relationship	Bladder Cervix Colon and rectum Esophagus Kidney Larynx Leukemia (AML) Liver Lung Oral cavity and pharynx Pancreas Stomach	4,983 447 N/A 8,592 3,043 3,009 1,192 N/A 125,522 4,893 6,683 2,484
Evidence Suggestive but Not Sufficient to Infer Causal Relationship	Breast	
Inadequate to Infer Presence or Absence of Causal Relationship	Ovary	
Evidence Sufficient to Infer No Causal Relationship	Prostate	



#### Oncogenic Viruses

- HPV (most commonly HPV-16) particularly in the oropharynx.
- tend to be younger
- less likely to have a strong history of tobacco and ethanol use,
- history of multiple sex partners,
- have a better prognosis, and
- appear to have a lower rate of SPTs.
- Association between Epstein-Barr virus and NPC.



#### Metabolism of Tobacco

• Nicotine is metabolized primarily to cotinine, which is further metabolized to trans-3'-hydroxycotinine (3HC), catalyzed by the liver cytochrome P450 2A6.



### **Smoking & Tar**

- Historically, tar was believed to be the main contributor to smoking-caused disease.
- Tar is not a specific substance, but simply the collected particulate matter from cigarette smoke, less water and nicotine.



## Pooled OR for chewing tobacco and risk of oral cancer = 4.7 [3.1–7.1]

Study or subgroup	log (odds ratio)	SE	Weight	Odds ratio IV, random, 95% CI	Year	Odds ratio IV, random, 95% CI
Nandakumar et al. 1990	2.55	0.27	6.6%	12.81 [7.54, 21.74]	1990	
Rao et al. 1994	1.29	0.21	6.9%	3.63 [2.41, 5.48]	1994	29 <del></del> 3
Khan et al. 1995	0.84	0.59	4.7%	2.32 [0.73, 7.36]	1995	<del></del>
Goud et al. 1990	2.14	0.33	6.3%	8.50 [4.45, 16.23]	1996	-
Wasnik et al. 1998	2.07	0.31	6.4%	7.92 [4.32, 14.55]	1998	<del></del>
Dikshit and Kanhere 20	00 1.75	0.24	6.8%	5.75 [3.60, 9.21]	2000	-
Znaor et al. 2003	1.61	0.08	7.4%	5.00 [4.28, 5.85]	2003	
Subapriya et al. 2007	1.12	0.18	7.1%	3.06 [2.15, 4.36]	2007	-
Gangane et al. 2007	2.3	0.2	7.0%	9.97 [6.74, 14.76]	2007	-
Basu et al. 2008	0.27	0.38	6.0%	2.05 [0.98, 4.33]	2008	•
Jayalekshmi et al. 2011	2.14	0.23	6.8%	8.50 [5.42, 13.34]	2009	-
Jayalekshmi et al. 2010	0.99	0.16	7.1%	2.69 [1.97, 3.68]	2010	*
Pednekar et al. 2011	0.18	0.09	7.3%	1.20 [1.00, 1.43]	2011	•
Madani et al. 2012	2.11	0.22	6.9%	8.25 [5.36, 12.69]	2012	
Ray et al. 2013	1.36	0.23	6.8%	3.90 [2.48, 6.12]	2013	-
Total (95% CI)			100.0%	4.72 [3.13, 7.11]		•
Heterogeneity: $\tau^2 = 0.59$	$\chi^2 = 266.77$ , df	= 14 (P	< 0.00001	); $I^2 = 95\%$	3	MILLANDER TREATMENT
Test for overall effect: Z =	= 7.41 (P < 0.000	01)			0.01	0.1 1 10 Manual

# Carcinogenic Agents in Smokeless Tobacco

- N-nitrosamines
- Benzene,
- 1,3-butadiene,
- Aromatic amines,
- Cadmium



# A dose of 10 gms of chewing tobacco for about 26 years was observed to have produced cancerous lesions in the buccal cavity.

Eur J Epidemiol. 1990 Jun;6(2):219-22.

Epidemiological correlates between consumption of Indian chewing tobacco and oral cancer.

Goud ML1, Mohapatra SC, Mohapatra P, Gaur SD, Pant GC, Knanna MN.

Author information

#### Abstract

The problem of cancer is universal; the only variation occurs in the type, site or other clinicoepidemiological parameters. Peculiarly enough, oral cancers caused by chewing tobacco are common in India and some parts of the Indian sub-continent. Oral cancers caused by other carcinogens are not common in these areas. The present study shows a significant association (P less than 0.001) between the use of Indian chewing tobacco and oral cancer. Number of quids, mean quantity of tobacco and mean duration of keeping the quids in the mouth had direct dose and effect relationships in causation of oral cancer. A dose of 10 gms of chewing tobacco for about 26 years was observed to have produced cancerous lesions in the buccal cavity.

# Pooled OR for Paan with tobacco and risk of oral cancer = 7.1 [4.5–11.1]

Journal of Cancer Epidemiol

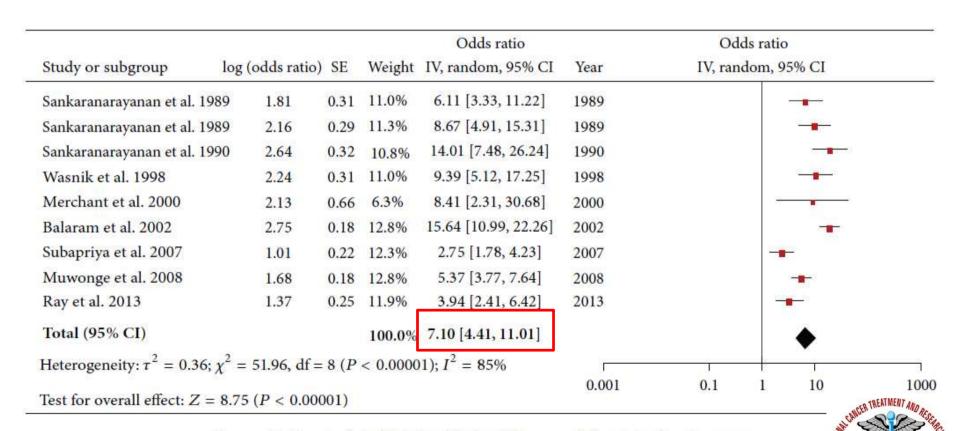


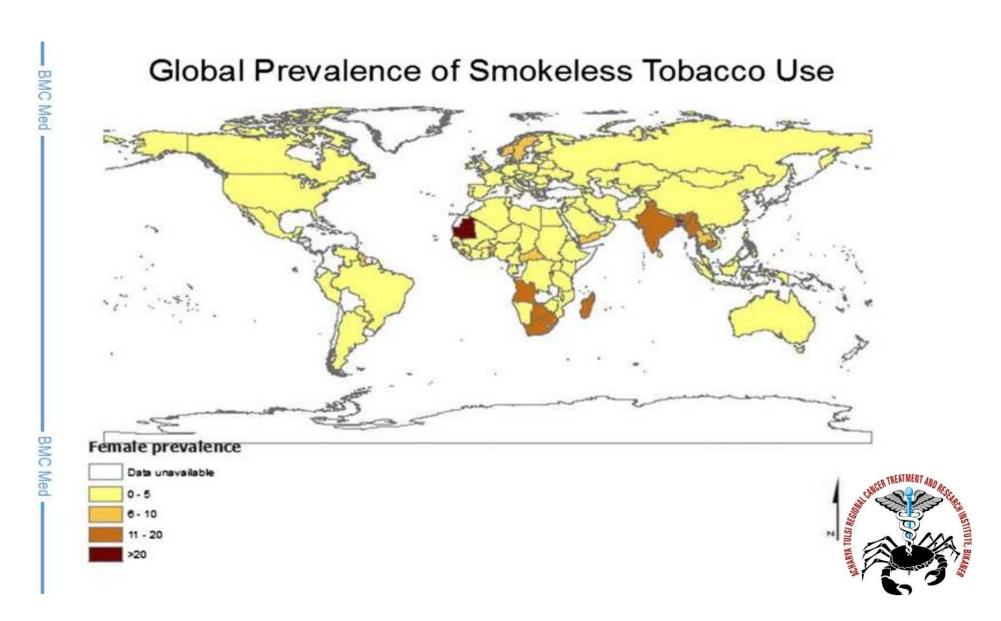
FIGURE 3: Forest plot of betel quid plus tobacco and the risk of oral cancer.

Ref: Zohaib Khan, et al. Smokeless Tobacco and Oral Cancer in South Asia: A Systematic Review with Meta-Analysis, Journal of Cancer Epidemiology, vol. 2014.

#### Hazard of Pan with Tobacco

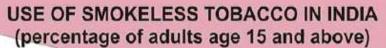
- Slaked lime: carcinogenic potential.
- Facilitates the production of reactive oxygen species in the saliva of chewers
- Also facilitates the hydrolysis of arecoline into arecaidine which in turn facilitates increased fibroblast proliferation and collagen synthesis, which are essential for premalignant changes.
- Areca nut in paan: carcinogenic properties

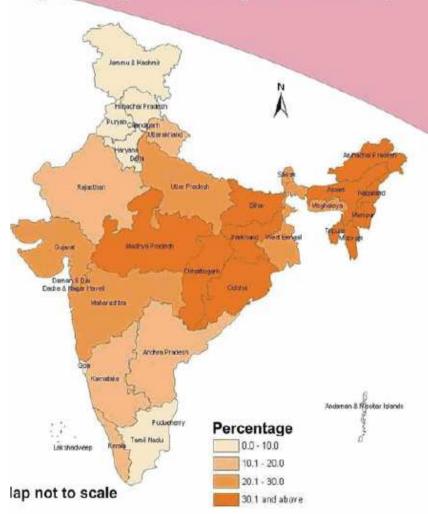
### An Important Point: SLT Use in Women





#### Report of GATS 2010



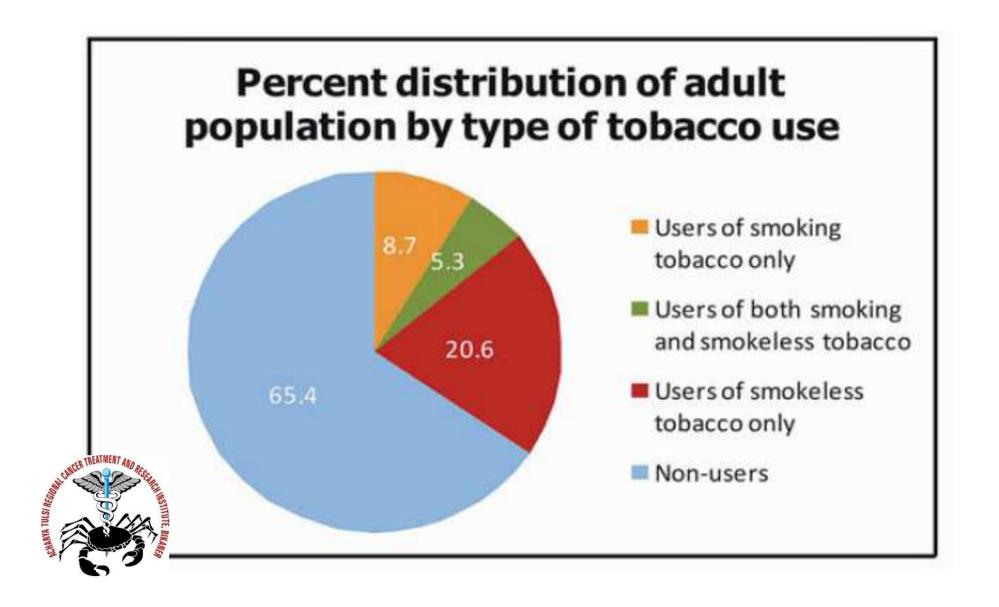




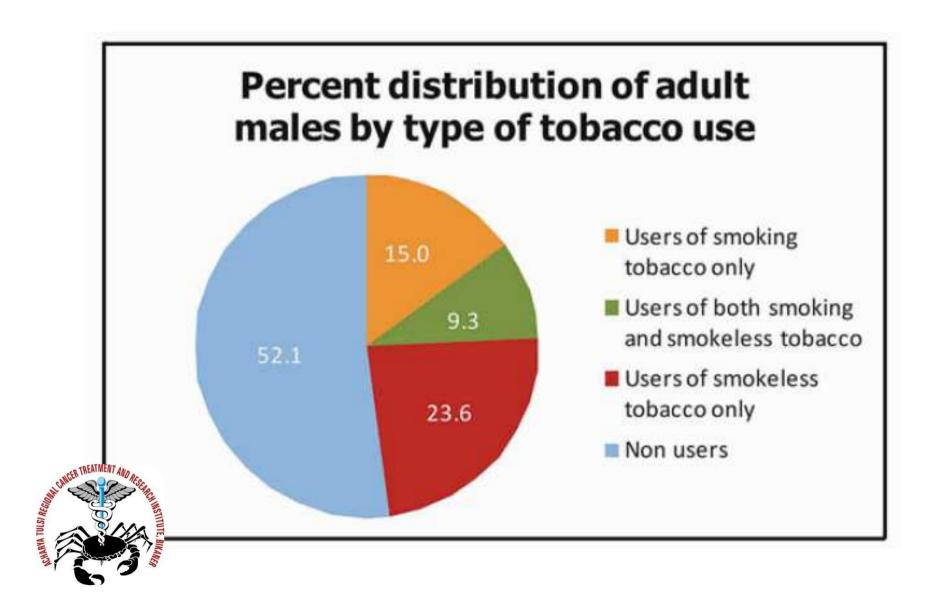
Ministry of Health and Family Welfare Government of India

Global Adult Tobacco Survey (GATS)

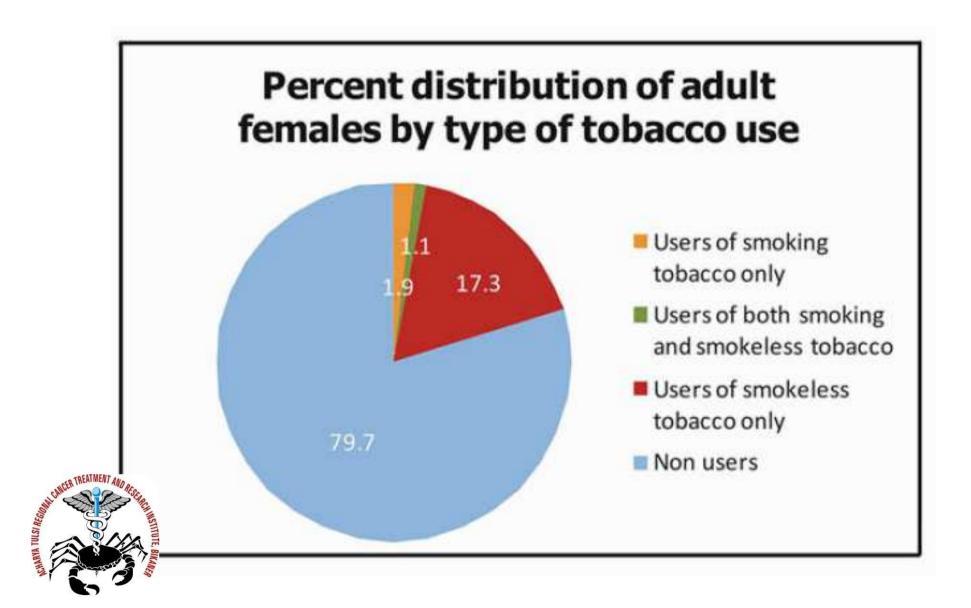
#### India: 34.6% adults consume Tobacco



#### India: 47.9% males consume Tobacco

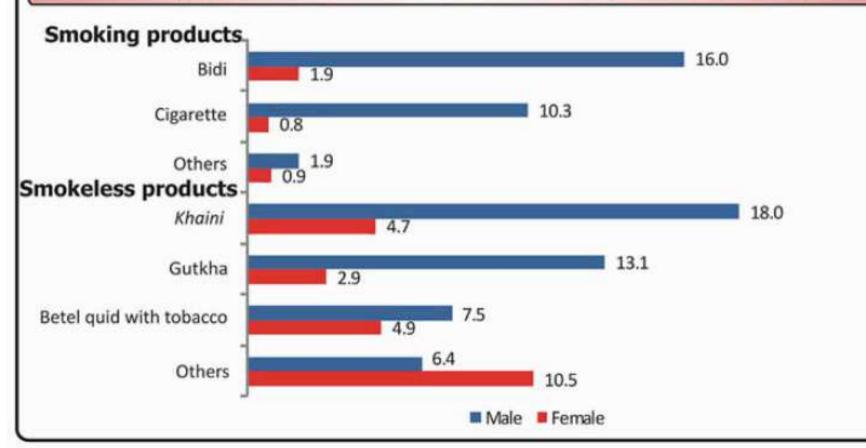


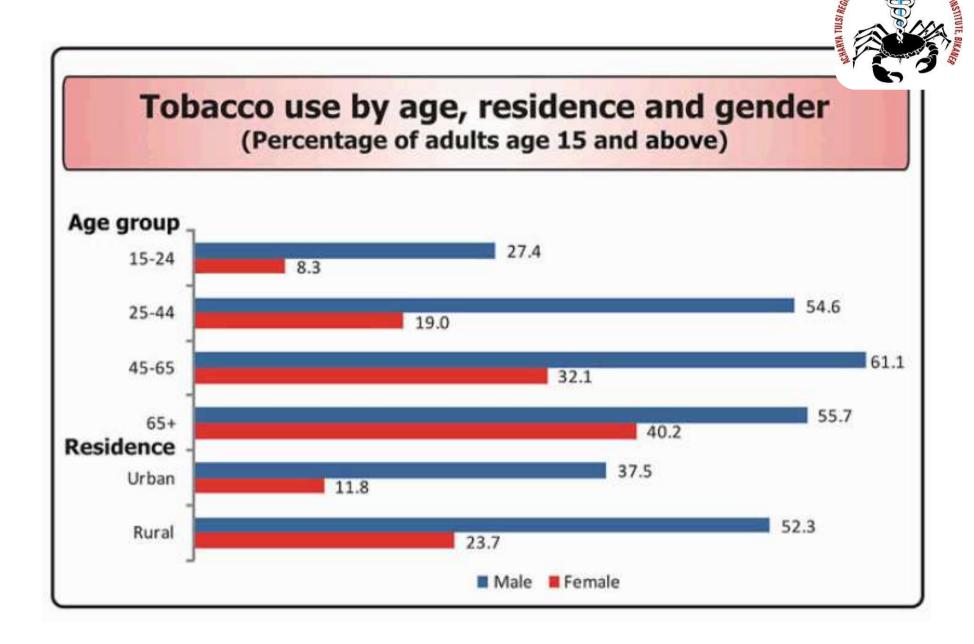
#### India: 20.3% females consume Tobacco











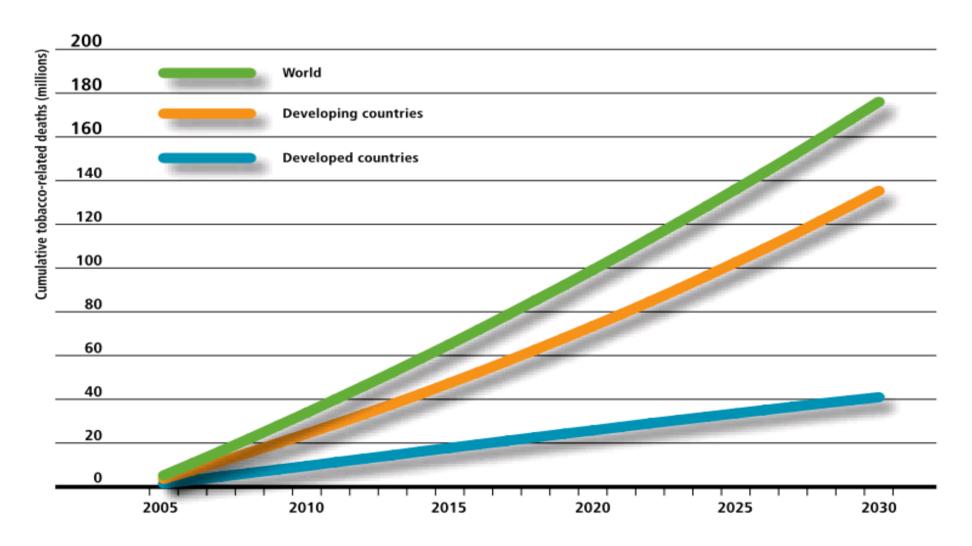
#### **Policy Implications**



- Most of the tobacco control initiatives: aimed towards cessation of smoking.
- Main strategy to decrease smoking prevalence: high amount of taxes levied on smoking products.
- This strategy may facilitate an unintentional push towards SLT use as its cheaper compared to smoking.
- Additionally, big tobacco companies revert to manufacturing SLT products and advertising them as less harmful than smoking.
- A surge in the use of SLT products and subsequent increased risks for oral cancer for the general public.
- Awareness of the potential dangers related to such approach and new programs for SLT cessation.

## TOBACCO WILL KILL OVER 175 MILLION PEOPLE WORLDWIDE BETWEEN NOW AND THE YEAR 2030

#### Cumulative tobacco-related deaths, 2005–2030



Source: Mathers CD, Loncar D. Projections of global mortality and burden of disease from 2002 to 2030. PLoS Medicine, 2006, 3(11):e442.



#### **SAY NO TO TOBACCO!**





## THANKS

