

Use of Non Cigarette Tobacco Products (NCTP)

Smokeless Tobacco Waterpipes Cigars Pipes

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NCTP Definitions & Products





Pipes









Cigars











Images from www.trinketsandtrash.org

Cigar Definition

U.S. Department of Treasury (1996):

Cigar



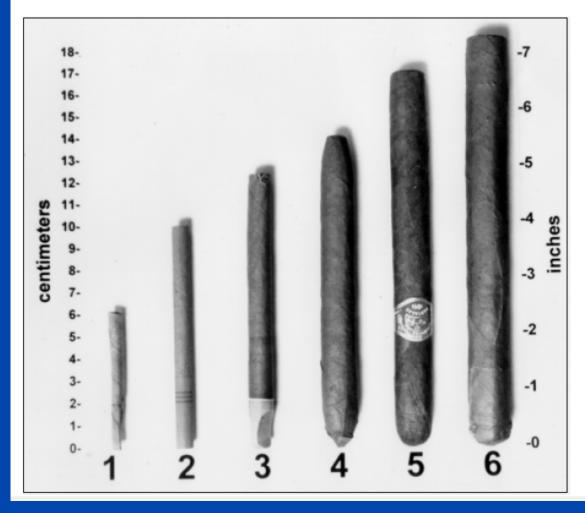
VS.

Cigarette

"Any roll of tobacco wrapped in paper or in any substance not containing tobacco."



Types of cigars on the U.S. Market in 1996: (1) bidi (imported from India), (2) little cigar with filter tip, (3) small cigar with plastic mouth piece, (4) regular cigar, (5) and (6) premium cigar.





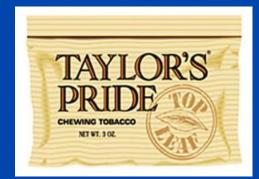


Rooster















Smokeless Tobacco

Chewing tobacco

- Loose leaf (i.e., Redman)
- Plugs
- Twists

Snuff

- Moist (i.e., Copenhagen, Skoal)
- Dry (i.e., Honest, Honey bee, Navy, Square)









"Chewing Tobacco" = Cut tobacco leaves



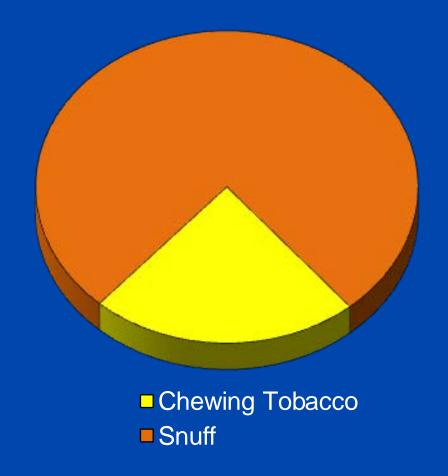


"Snuff" = Moist ground tobacco





Type of ST Used in U.S.





National Survey on Drug Use and Health (NSDUH)

"Spitless Tobacco" - Star Scientific







RJ Reynold's







"Swedish Style" ST





Phillip Morris (Altria)





"Fully Dissolvables"





Tobacco-less Nicotine Product - Altria

Package of 16 discs, each containing about 1.5 milligrams of nicotine = \$3

Virginia test market





Waterpipe

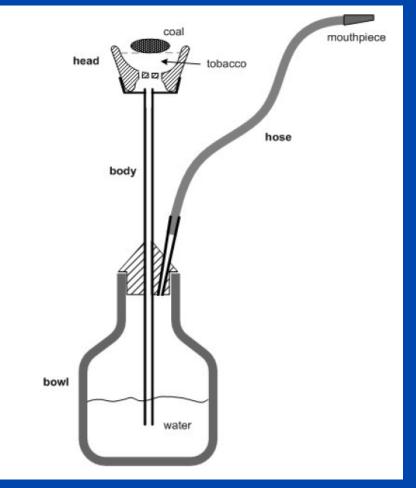
- Use dates back 4 centuries in Asia & Africa
- Typically consumed socially with friends and family
- Becoming trendy in US among youth
- Hookah "cafes" gaining popularity
- Smoke described as smoother and more flavorful
- Often (mis)perceived as healthier alternative due to water filtration, cooler mouth feel, and reduced irritation





Anatomy of a Waterpipe







Types of Waterpipe Tobacco

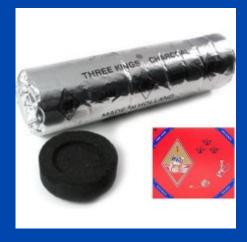
- Maasel/Mu'essel
 Combination of tobacco and molasses, honey or fruit
- Tumbak/Ajami
 Dark tobacco paste
- Jurak
 Combination of tobacco and fruits, oils, honey or molasses. May be flavored or flavorless
- Moist tobacco

 requires charcoal to keep burning



Hookah tobacco

Charcoal





Source: Knishkowy & Amitai. (2005). Pediatrics, 116, e113-e119.

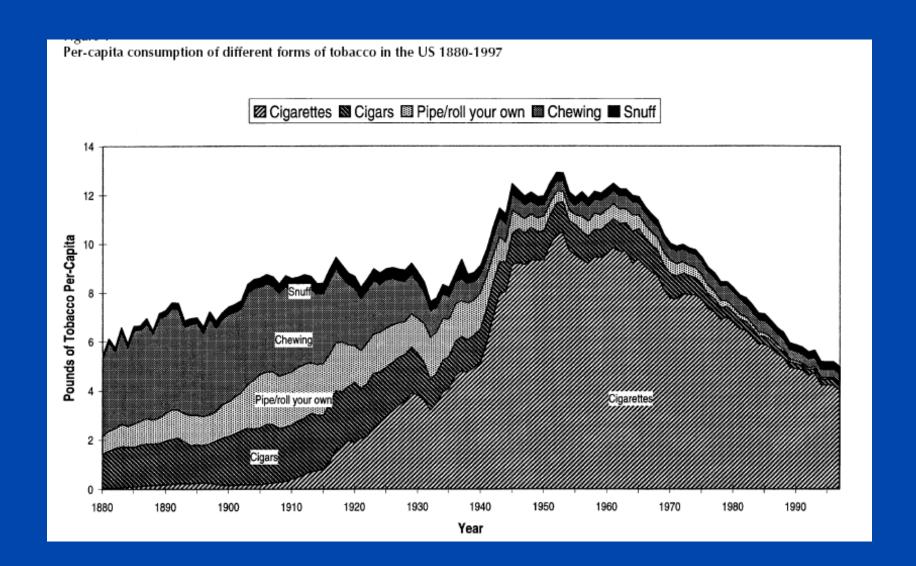




NCTP: Trends & Prevalence

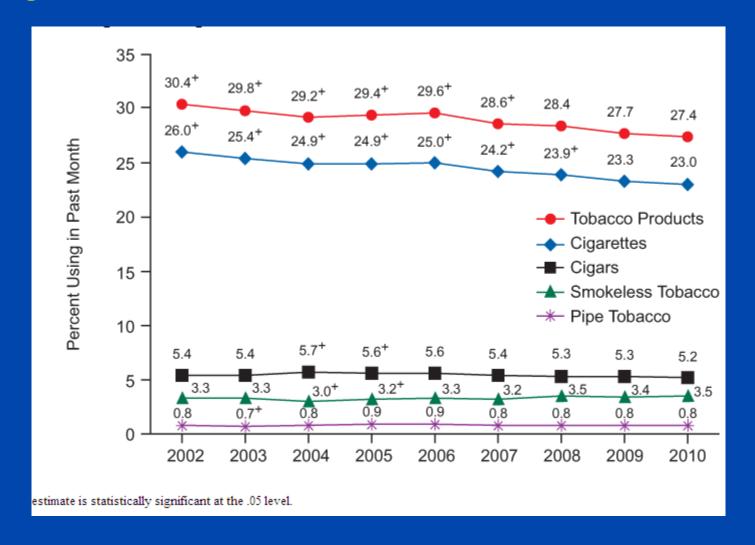








Past Month Tobacco Use among Persons Aged 12 or Older

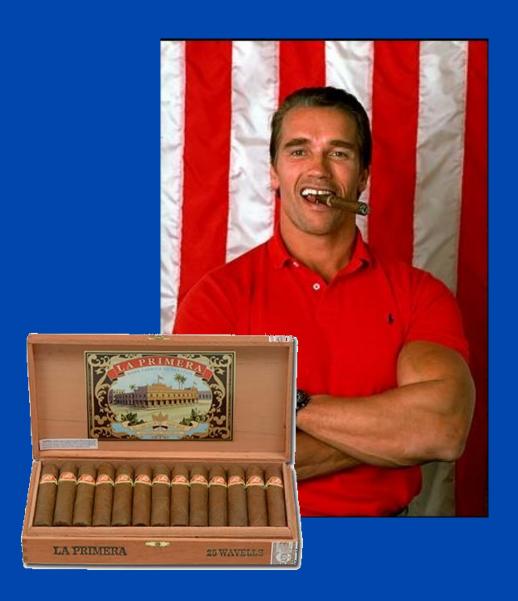








Cigar Smoking



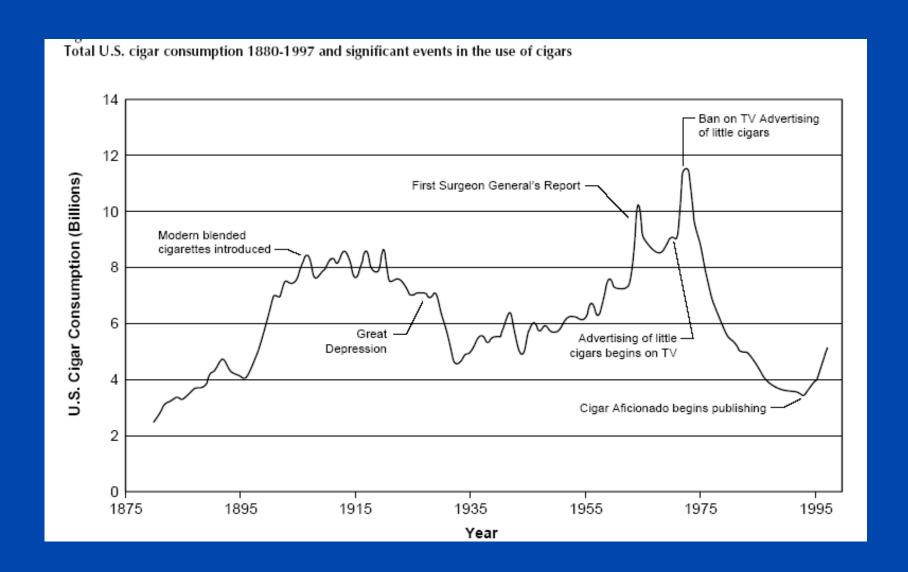


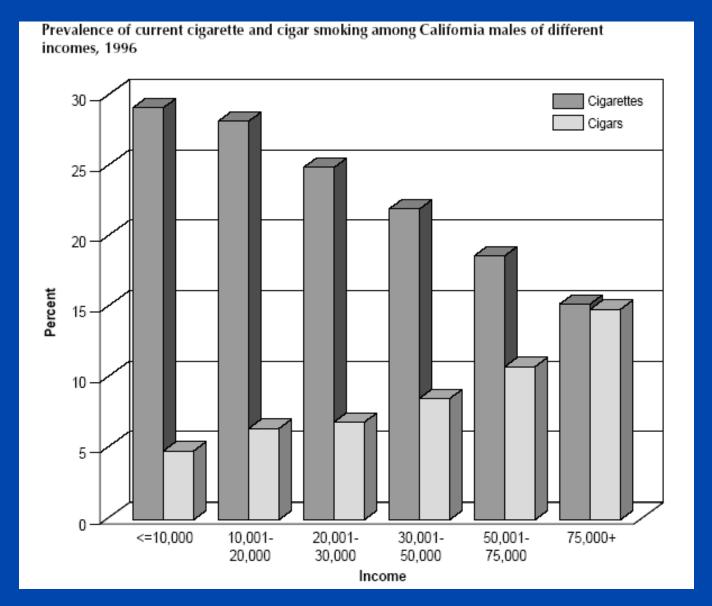


Figure 6.4 Annual Numbers of New Users of Tobacco: 1965–2001 4,500 4.000 ousands of New Users 3,500 3,000 2,500 2,000 Daily 1,500 igarettes 1 000 Tobacco 500 1965 1970 1975 1980 1985 1990 1995 2000

< 18 years of age group constituted an increasingly greater proportion of the number of new cigar smokers:

SAMHSA, Office of Applied Studies, National Survey on Drug Use and Health, 2002. The Monitoring the Future Study, University of Michigan, 2001 and 2002.





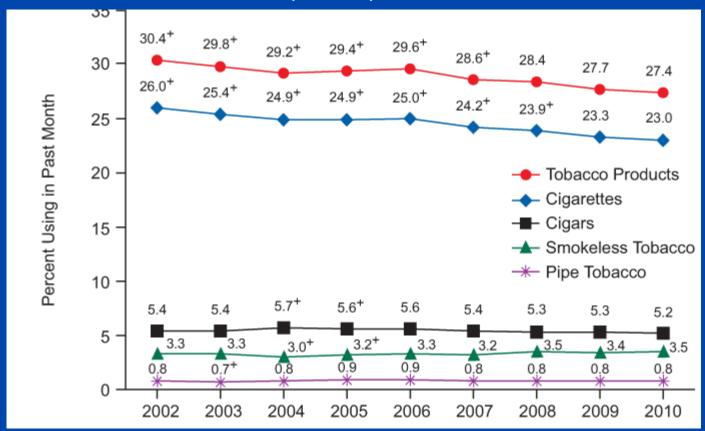


NCI Monograph 9. Cigars: Health Effects and Trends.



Prevalence of ST Use

In 2010, 8.9 million (3.5%) used smokeless tobacco





SAMSHA

Percentages of
Persons Aged 12 or
Older Using
Smokeless Tobacco in
the Past Month, by
Demographic and
Geographic
Characteristics: 2002
to 2007



Demographic and Geographic Characteristic	Past Month
Age Group in Years	
12 to 17	2.2%
18 to 25	5.0%
26 or Older	3.0%
Gender	
Male	6.2%
Female	0.4%
Race/Ethnicity	
White	4.1%
Black or African American	1.4%
American Indian or Alaska Native	7.1%
Native Hawaiian or Other Pacific Islander	2.9%
Asian	0.8%
Hispanic or Latino	0.9%
Two or More Races	2.9%
County Type	
Large Metropolitan	1.9%
Small Metropolitan	3.7%
250,000 to 1 Million Population	3.2%
<250,000 Population	4.7%
Non-Metropolitan	6.6%
Urbanized	5.5%
Less Urbanized	7.1%
Completely Rural	8.4%
Region	
Northeast	1.7%
Midwest	3.7%
South	4.2%
West	2.4%



Waterpipe

- After cigarettes, waterpipe use is the most common form of tobaccouse among university students
- Predominantly young, males
- 30% ever use & 8.4% current use
- Used in a social context
 - More common in fraternities/sororities
- Most smoked < 2 years
 - 10% daily
- Most perceive less addictive and harmful
 - 67% said "cigarettes more harmful"





Odds of Trying Waterpipe, Snus, or ENDDs (n = 3158)

Predictors	Have tried one of these products adjusted OR (95% confidence interval)	
Smoking status		
Former smoker	2.71 (2.06, 3.56)	
Nondaily smoker	6.13 (4.02, 9.33)	
Daily smoker	5.53 (4.03, 7.58)	
Region		
Northeast	1.68 (1.16, 2.42)	
Midwest	1.65 (1.20, 2.28)	
West	1.80 (1.36, 2.39)	
Age		
18–24	2.18 (1.60, 2.97)	
Sex		
Males	3.51 (2.77, 4.45)	
Education		
High school	1.58 (.99, 2.51)	
Some college	2.67 (1.69, 4.22)	
College degree	2.04 (1.26, 3.30)	

*Lifetime Use

McMillen R, et al. Use of emerging tobacco products in the United Model also included race, not significant. Reference groups were as follows: States. J Environ Public Health. never smokers, south region, 25 years of age and older, females, and no high 2012;2012:989474.



school degree.



Health Impact of NCTP



Adjusted relative risk* (95% CI) of death Sustainer = No switching



	'Sustainers'		
Deaths from	Never smoked	Pipe only	Cigarettes only
Men (n=16 932 aged 20-49 years)			
All causes	1.00	1.99 (1.73 to 2.27)	2.44 (2.27 to 2.62)
Ischaemic heart disease	1.00	3.07 (2.35 to 4.00)	3.17 (2.69 to 3.73)
Stroke	1.00	1.54 (0.84 to 2.82)	2.30 (1.71 to 3.11)
Cardiovascular disease	1.00	2.49 (1.99 to 3.10)	2.81 (2.48 to 3.20)
Lung cancer	1.00	10.32 (5.55 to 19.18)	16.78 (10.31 to 27.33)
Other smoking related cancer	1.00	1.47 (0.99 to 2.18)	1.95 (1.59 to 2.38)



Disease Risks

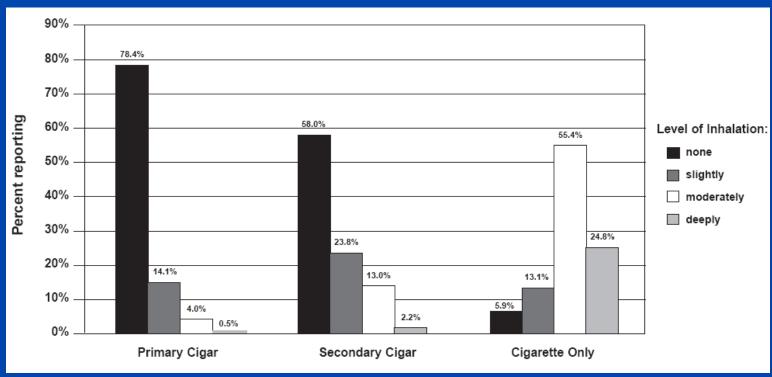


- Tar of cigar is more carcinogenic than cigarette smoke tar
- Morbidity and mortality correlates with:
 - +/- inhalation
 - depth of inhalation
 - number of cigars they smoke



Levels of Inhalation, CPS-1 Study



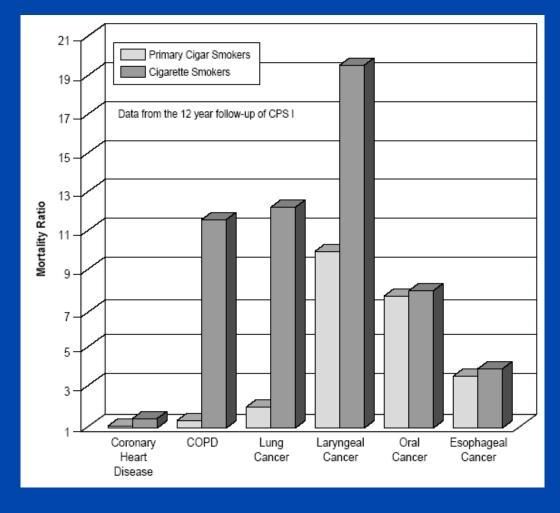




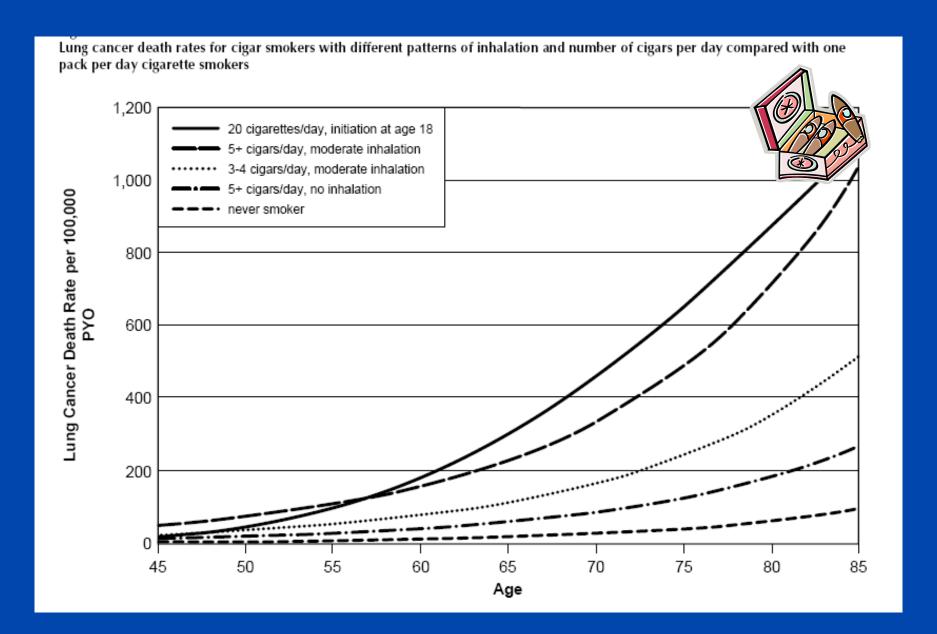
NCI Monograph 9. Cigars: Health Effects and Trends.

Mortality Ratios for Cigar & Cigarette Smokers vs. Never Smokers





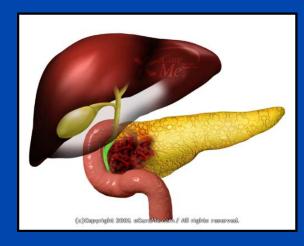






Cigar Smoking – Pancreatic Cancer Risk

- Compared with never tobacco users, cigar-only smokers have an increased risk for pancreatic cancer
 - OR 1.6 (95% CI: 1.2-2.3)
- Comparable to that of cigarette-only smokers
 - OR 1.5 (95% CI 1.4-1.6)





ST – Health Consequences

Report on Carcinogens, 10th Edition, National Toxicology Program, USDHHS

Smokeless tobacco

"Known to be a human carcinogen"









- Including.....
- β-Angelica lactone
- Coumarin
- Ethyl carbamate (urethane)
- Formaldehyde
- Acetaldehyde
- Crotonaldehyde

Smokeless Tobacco and Some Tobacco-specific N-Nitrosamines. IARC Working Group on the Evaluation of Carcinogenic Risks to Humans (2004)

- Tobacco-specific N-nitrosamines (TSNA)
 - N'-Nitrosonornicotine (NNN)
 - 4-(Methylnitrosamino)-1-(3pyridyl)-1-butanone (NNK)
 - 4-(Methylnitrosamino)-1-(3pyridyl)-1-butanol (NNAL)
 - N'-Nitrosoanabasine (NAB)
- Arsenic
- Nickel compounds
- Polonium-210
- Uranium-235
- Uranium-238



Health Effects: Cancers - U.S. Data

Location OR (95%	6 CI)
------------------	-------

Cancer, Mouth and Gum 11.2 (4.1-30.7)^A

Gum & Buccal Mucosa 4.2 (2.6-6.7)

Larynx 7.3 (2.9-18.3)^A

Salivary gland 5.3 (1.2-23.4)^A

Kidney 4.0 (1.2-12.9)^C

Pancreatic 3.6 (1.0-12.8)^D

- A Stockwell HG, et al. Head Neck Surg. 1986 Nov-Dec;9(2):104-10.
- B Winn DM, et al. N Engl J Med. 1981 Mar 26;304(13):745-9.
- C Goodman MT, et al. Am J Epidemiol. 1986 Dec;124(6):926-41.
- D Muscat JE, et al. Cancer Epidemiol Biomarkers Prev. 1997 Jan; 6(1):15-9.



ST Health Effects: CV Disease

Rooste

CPS-II

- *Current ST use vs. never associated with death from:
 - All causes: HR 1.18 (95% CI: 1.08-1.29)
 - CHD: HR 1.26 (95% CI: 1.08-1.47)
 - Cerebrovascular dz: HR 1.40 (95% CI: 1.10-1.79)
- No difference between snuff and chewing tobacco
- Former use did increase the risk of death in any category

*Multivariate-adjusted



ST – Oral Lesions

- Leukoplakia
- Oral cancer
- Dental disease
 - erosion of enamel
 - dental caries
- Periodontal Disease
 - gingival recession
 - soft tissue/hard tissue loss
 - gingivitis





Waterpipe – Health Effects

- 1-hour session involves inhaling 100-200 times volume of smoke from a single cigarette
- Smoke contains CO, heavy metals, and carcinogens
- Charcoal added to keep tobacco burning increases health risks
- Sharing = tuberculosis & hepatitis

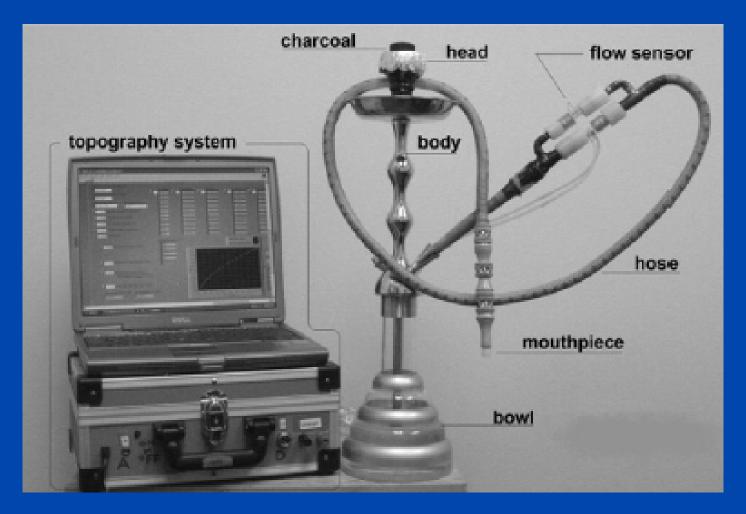
WHO. TobReg Advisory Note. Waterpipe Tobacco Smoking

Google: "who tobreg water pipe"





Waterpipe Analysis





Eissenberg T, et al. American journal of preventive medicine. 2009 Dec;37(6):518-23.

Waterpipe vs. Cigarette

- Directly compare waterpipe use & cigarette smoking
- 31 participants reporting monthly waterpipe use & weekly cigarette smoking



- Cross-over: 45 minutes with waterpipe (WP) and 1 cigarette
- CO (carbon monoxide): 23.9 ppm WP vs. 2.7 C
- COHb (carboxyhemoglobin): 3.9% WP vs. 1.3% C
- Puff volume: 48.6 L WP vs. 1.0 L C
- Peak nicotine levels were comparable
 - 1.7 times the nicotine exposure



Disease Risks

- Burning charcoal is normally placed atop the tobacco to smoke the narghile waterpipe
- Waterpipe smokers thus also inhale large quantities of combustion-generated toxicants





Waterpipe – Health Effects

Single narghile smoking session:

- 50 times the quantities of carcinogens as one cigarette^A
- Many times the formaldehyde, acetaldehyde and acrolein typically found in a cigarette^B
- 2.25 mg nicotine^C
- 242 mg nicotine-free dry particulate matter (NFDPM)^C
- Higher levels of arsenic, chromium and lead than a cigarette^C
 - A Sepetdjian E, et al. Food Chem Toxicol. 2008 May;46(5):1582-90.
 - B Al Rashidi et al. Volatile aldehydes in the mainstream smoke of the narghile waterpipe. Food Chem Toxicol. 2008 Nov;46(11):3546-9.
 - C Shihadeh A. Investigation of mainstream smoke aerosol of the argileh water pipe. Food Chem Toxicol. 2003 Jan;41(1):143-52.



Waterpipe

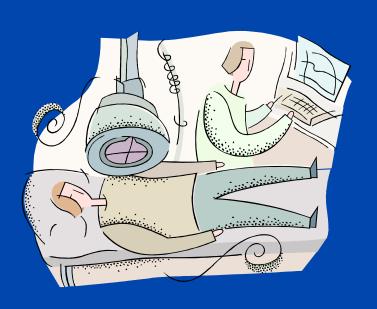
- Waterpipe tobacco smoking negatively affects lung function
 - Significant reduction in FEV1 compared to no smoking (4% lower FEV1)
 - Trend toward lower FVC (1.38% lower FVC) compared to no smoking
 - No statistically significant difference in FEV(1), FVC, and FEV(1)/FVC compared to cigarette smoking
- May be as harmful as cigarette smoking
- Likely to be a cause of COPD





Waterpipe

- Literature review
 - Cohort, case-control and cross-sectional studies
- Increased risk for
 - Lung cancer
 - Respiratory illness
 - Low birth-weight
 - Periodontal disease







Pharmacology



NCTP Bioavailability of Nicotine (aka "Smoke Yields")

<u>Type</u>		Nicotine (mg)	
Cigar	ette (filter)	1.1	
Pipe		5.2	
Smok	keless tobacco		
	Chewing tobacco	4.5	
	Moist snuff	3.6	
Cigar	S		
	Little cigars, Swishers	3.8	
	Premium, Macanudo	13.3	
4 mg	nicotine gum	1.9	



Smokeless Tobacco Nicotine "Content"

- 4.8 mg nicotine/gm of moist snuff x 30 gm/can = 144 mg
- 144 mg nicotine/(1.8 mg nicotine/cigarette) =
 80 cigarettes
- 80 cigarettes/(20 cigarettes/pack) = 4 packs
- 1 can snuff = 4 packs of cigarettes
- ST Users are exposed to as much, and possibly more, daily nicotine than cigarette smokers













Table 1. Tobacco-specific nitrosamines, pH, total and unprotonated nicotine, and minor tobacco alkaloids in smokeless tobacco products.

						_	Alkaloids, mg/g dry weight							
	TSNAs ^a , μg/g dry weight						Nicot	ine						
Product	NNNª	NNKª	NATa	NAB ^a	Total	pН	Total	Free	Nornicotine	Anatabine	Anabasine			
New products														
Taboka														
Original	1.05	0.077	0.370	ND^b	1.50	6.64	21.1	0.844	1.04	3.78	0.149			
Green	0.948	0.092	0.292	0.002	1.33	6.85	19.9	1.26	1.02	4.03	0.197			
Marlboro Snus														
Rich	1.27	0.259	0.455	ND	1.98	6.83	17.8	1.08	0.438	2.60	0.111			
Mild	1.52	0.229	0.234	ND	1.98	6.47	12.8	0.350	0.484	1.82	0.072			
Spice	1.56			2.06	6.85	17.9	1.13	0.411	2.17	0.097				
Mint	3.28	0.215	0.221	ND	3.72	6.58	20.0	0.701	0.454	1.97	0.063			
Camel Snus														
Original	1.15	0.270	0.297	0.012	1.73	7.46	28.2	6.09	0.353	1.39	0.164			
Spice	1.27	0.157	0.305	0.015	1.75	7.75	25.4	9.16	0.314	1.09	0.183			
Frost	1.20	0.267	0.204	0.009	1.68	7.59	23.7	6.40	0.313	0.741	0.103			
Skoal Dry			=.											
Regular	3.57	0.360	0.478	ND	4.41	7.23	11.3	1.57	0.345	1.41	0.117			
Cinnamon	5.30	0.313	0.572	0.002	6.19	6.85	11.9	0.751	0.324	1.02	0.130			
Menthol	2.53	0.279	0.203	ND	3.01	7.18	11.9	1.51	0.386	1.37	0.127			
Mean for new products	2.05	0.231	0.323	0.008	2.61		18.5	2.57	0.490	1.95	0.126			
Traditional products														
General Snus	1.66	0.464	0.969	0.008	3.10	7.95	16.7	7.69	0.223	0.367	0.072			
Copenhagen Snuff	5.12	1.40	1.12	0.152	7.79	7.45	23.0	4.88	0.248	1.43	0.150			
Copenhagen Long Cut	3.76	1.10	1.35	0.062	6.27	7.53	26.7	7.14	0.157	0.770	0.037			
Skoal Long Cut	4.66	1.64	1.59	0.074	7.96	7.51	25.6	6.03	0.233	1.02	0.049			
Kodiak Wintergreen	6.86	1.41	3.58	0.179	12.0	8.23	19.6	12.1	0.164	0.438	0.055			
Mean for traditional products	4.41	1.20	1.72	0.095	7.42		22.3	7.57	0.205	0.805	0.073			

Note. ^aAbbreviations: TSNAs, tobacco-specific N-nitrosamines; NNN, N'-nitrosonornicotine; NNK, 4-(methylnitrosamino)-1-(3-pyridyl)-1-butanone; NAT, N'-nitrosoanatabine; NAB, N'-nitrosoanabasine. ^bND, not detected.



ST Characteristics Affecting Nicotine Blood Concentrations

- Concentration of nicotine in ST product
- Size of the tobacco cuttings
 - Long cut
 - Fine cut (higher)
- Ammonium bicarbonate (additive)
 - Lower acid level of product = higher free nicotine
- Acetic acid (additive)
 - Increases salivation enhances absorption





pH Manipulation by Industry

- October 1994, Wall Street Journal:
- "US Tobacco routinely adds chemicals to its snuff to deliver the free nicotine faster and to make the product stronger."
 - Larry Story, former UST chemist
- "It (Copenhagen) was brought up to a pH of 7.8 by adding more sodium bicarbonate and ammonium carbonate."
 - Larry Story, former UST chemist26



Impact of pH Manipulation: Long-Term

- Likelihood of choosing a brand with higher nicotine content is related to:
 - Increasing duration of use
 - Increasing intensity of use
 - Frequency of ST use
- ST users who have used higher nicotine-containing products are more likely to report:
 - More nicotine withdrawal symptoms
 - Continued use because of <u>difficulty quitting</u>



Waterpipe

- Data indicates that daily waterpipe use of the produced a 24-hr urinary cotinine level of: 0.785 microg/ml (95% CI = 0.578-0.991 microg/ml)
- Daily waterpipe smoking is equivalent to smoking 10 cigarettes (95% CI: 7-13)







Current Recommendations for Treatment

Pipes, Cigars, & Waterpipe









Treatment Options

Non-daily users

- Nicotine gum
- Nicotine lozenge
- Nicotine inhaler
- Nicotine nasal spray

Daily users

- Nicotine patch
- Bupropion SR
- Varenicline







Smokeless Tobacco (ST)



Assessing Dependence in ST Users

Number of cans per week

- Strongest correlation with nicotine/cotinine blood concentrations
- Used for dosing guidelines





Nicotine Patch Dosing Algorithm for ST Users

	Peak serum nicotine concentrations (ng/mL)	Cans or pouches per week	Patch dose
Low	0-10	< 2	14 mg/d
Intermediate	11-20	2-3	21mg/d
High	> 20	> 3	42 mg/d



Ebbert. JSAT. 2004



Bupropion SR: Dosing

150 mg daily for 3-4 days then



150 mg twice a day for 3-4 days

THEN

STOP CHEWING



3 to 12 months - No taper needed





Snuff Substitutes

- Smokey Mountain®
- Golden Eagle®
- Oregon Mint®
- KIK IT®
- Jerky®
- Bacc-Off®



















Nicotine Lozenge

2 mg & 4 mg

Dissolves in mouth over 20-30 minutes

 Delivers 25% more nicotine than the gum







Nicorette "Mini" – 2 mg/4 mg

Generic (large) lozenge





Nicotine Lozenge: Dosing

- Not to be chewed or swallowed whole
- Avoid eating or drinking food during and 15 minutes prior to use
- Monotherapy
 - 2 mg
 - First dip \geq 30 min
 - 4 mg
 - First dip < 30 min
 - > 3 cans/week
- Combination may be optimal (patch)
- 1-2 lozenges every 1-2 hours
- Minimum of 9/day
- Taper over 12 weeks





Nicotine Gum

- Monotherapy
 - 2 mg
 - First dip \geq 30 min
 - < 2 cans/week</p>
 - 4 mg
 - First dip < 30 min
 - > 2 cans/week
- "Chew and Park"
- Combination with nicotine patch may be optimal





Scandinavian Snus



32% of men aged 16-35 use snus daily



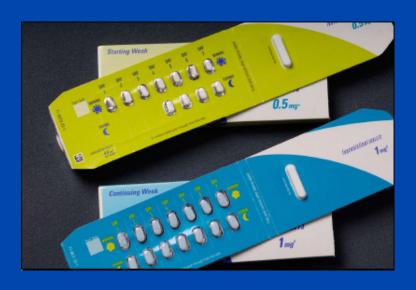
19% adult snus use prevalence



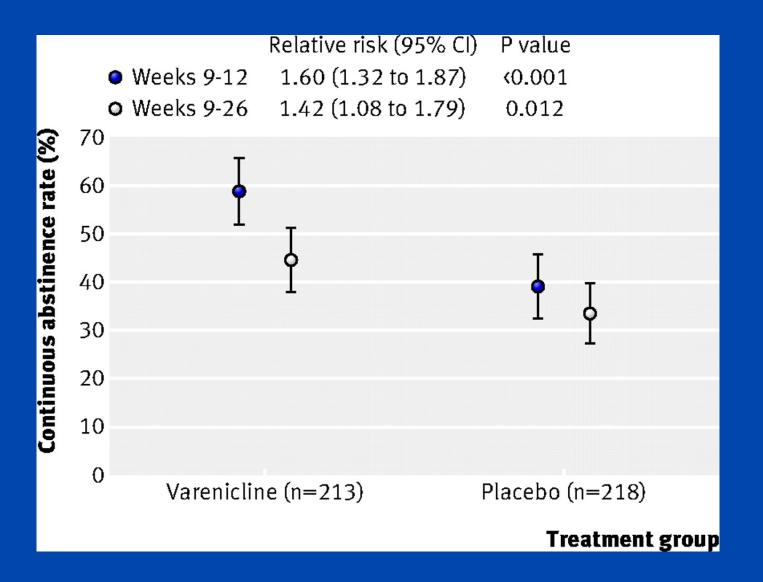


Varenicline (Chantix[™]) for Snus Users

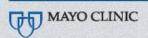
- Norway (7 sites) & Sweden (9 sites)
- Male/female daily ST users
 - Use at least 8 times/day
- Randomized to:
 - Varenicline for 12 weeks
 - Placebo
- Biochemical confirmation of abstinence
 - Salivary cotinine > 15 ng/mL



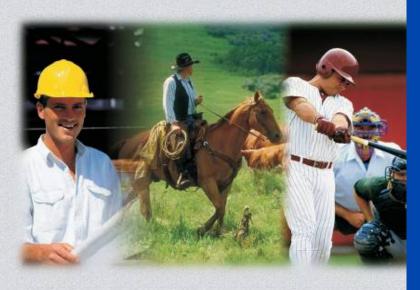








Your Path to Smokeless Tobacco Freedom

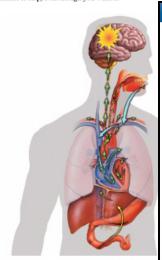


MAYO CLINIC NICOTINE DEPENDENCE CENTER



Your brain and nicotine: The physical challenge of quitting

You use chew for many reasons. One main reason is because smokeless tobacco contains an addicting drug called nicotine. This is the substance that makes it so difficult to stop, even though you want to.



Nicotine from chewing tobacco is absorbed into your bloodstree mouth (blue arrows). Nicotine from swallowed tobacco juice is al small intestine and then passes through your liver before enterin arrows). Your heart pumps nicotine to your brain and the rest of

You're prepared! It's your quit day!

It's the start of your tobacco-free life

My Quit Date is:

In preparation for today, I will:

- Clean out my car
- ☐ Make plans with my support people
- ☐ Plan activities or projects
- ☐ Dispose of all tobacco (check my gym bag, glove compartment and other storage areas)
- ☐ Make an appointment to se clean my teeth
- ☐ Make an appointment to se
- prescriptions

 Buy the medications recomm
- Other:

You may find it check mark next



CHEW CHECKS: We pitters "these cheeks" anothed to your timor pouch within rutberband. Remove these cards by kilding and tearing along the period sed lines.

Carry those with you and record each time you take a cheek — see page 17 for instructions.

_						D.A	TR:				DATE:							
Time of day	dlp/	Add more obser		Mo-od One word (angry, sad happy, contient)	Activity What you were doing at the time	Ma	Time of day	Mirw dip/ abov	Add more obser	Maed S = ritong N = recdepile L = light	Mood One word (argry, sad happy, content)	Activity What you were doing at the time	*	a. Three of day	Meer dip/ aheer	Add more obser	Mead Sentrong Nemodesale Lelight	Mood One to (argr) happy
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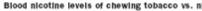
Bupropion Available dose: 150 mg

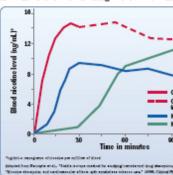
Comments:



Varenicline Available doses: 0.5 mg 1 mg

Comments:





Nicotine in chewing tobacco is absorbed very rapidly and peaks within placement. Even after the chew is removed, nicotine is absorbed through and small intestine. In comparison, the nicotine absorbed through the of the mouth (from gum) is absorbed much more slowly and often residualized with chewing tobacco.



Normal inside cheek





Snuff dipper pouch (where chew is placed)





Recommended ST Treatment Approach

- Bupropion SR
 - Weight gain prevention
 - Craving reduction
- Tailored nicotine patch therapy
 - Craving reduction
 - Short-term (end-of-treatment) abstinence
- Nicotine lozenge (short-term abstinence)
- Nicotine gum (craving reduction)
- Varenicline









Treatment Not Recommended for ST Users

- Nicotine inhaler
 - Designed to replicate the tactile sensation of a cigarette

- Nicotine nasal spray
 - Speed of intranasal delivery designed to the speed of delivery of a cigarette



Goals & Objectives

Review NCTP definitions & products



- Discuss prevalence/trends of NCTP
- Describe NCTP pharmacology
- Discuss NCTP dependence measures/withdrawal
- Review recommended treatments for NCTP

