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# TOBACCO PLANT FAMILY: SOLANACEAE GENUS: NICOTIANA RUSTICA





Rank	Country	Production (Million Tonnes)
1.	China	3.400
2.	India	.875
3.	Brazil	.810
4.	USA	.345
5.	Indonesia	.226
6.	Malawi	.151
7.	Argentina	.148
8.	Tanzania	.120
9	Zimbabwe	.115
10.	Pakistan	.102
•	TOTAL	6.333



# CONSUMABLE TOBACCO PRODUCTS

 SMOKELESS products used for: Chewing, Snuffing & Local application

Around 4000 chemicals found in smokeless tobacco (tobacco leaf).

SMOKING is in

#### Form of:

- Cigarette/Beedi
- > Chillum/Hukkah
- Cigar/Cheroot

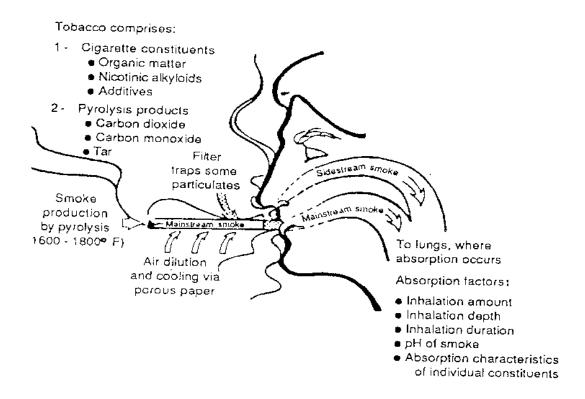






# **TOBACCO SMOKING**

 Lighted cigarette at 800oC gives the smoker, with each puff, hot potpourri of gases & many sized particles from main-stream smoke & side-stream smoke.





# CONSTITUENTS OF TOBACCO SMOKE



#### Many chemically active & most damaging

- Nicotine main chemical, highly addictive, causes many patho-physiological changes in the body.
- Tar suspended particulate matter & carcinogen.
- Carbon monoxide toxic gas, reduces oxygen carrying capacity of Hb.
- Nitrogen oxide damages lung tissue causing emphysema.
- Hydrogen cyanide allows toxic smoke ingredients accumulate in lung tissue.
- Ammonias damage lung tissue.
- Metals like arsenic, cadmium & lead. Some of these are carcinogenic.
- Radioactive compounds are carcinogenic.



# CARCINOGENS IN TOBACCO SMOKE

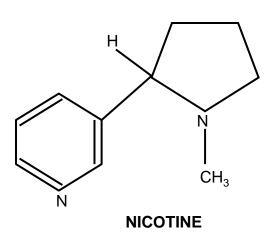
#### 70 carcinogens identified.

- N-Nitrosamines cause DNA mutation. NNK (Nicotine-derived nitrosamine ketone) & NNN (nitrosonornicotine)
- Benzene causes chromosaomal changes in bone marrow leading to anaemia
- Aromatic amines: high in side-stream smoke, confirmed human carcinogen.
- Aldehydes: Formaldehyde human carcinogen & acetaldehyde animal carcinogen
- □ 1,3-Butadine is human carcinogen & teratogen.
- Acrolein unsaturated aldehyde associated with lung cancer & DNA mutagen.
- Polyaromatic hydrocarbons, are carcinogenic, mutagenic & teratogenic. High PAH exposure in pregnant womb leads to low IQ in the child.

# **NICOTINE IN TOBACCO**

- □ In pure form:
  - Colourless, volatile, alkaline liquid,
  - Turns pale yellow to dark brown on exposure to air giving tobacco smell.
- Isolated from leaves by Posselt & Reimanbasic in 1828.
- Highly toxic & lethal causing pathophysiological changes
- One-drop sufficient to kill dog (or a man) within minutes.
- In smokeless tobacco dissolved in moisture of tobacco leaf as water soluble salt,
- In burning cigarette volatilizes & suspended on tar droplets.

#### An alkaloid



1-methyl-2-[3-pyrodyl] pyrrolidine  $C_{10}$ ,  $H_{14}$  &  $N_2$  double ring structure

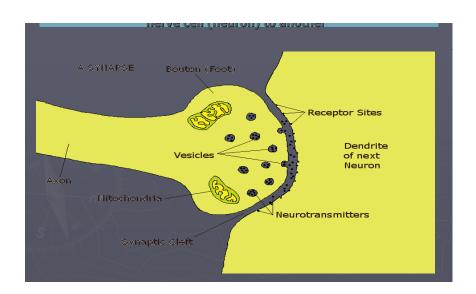


# **ABSORPTION OF NICOTINE**

- Conveyed through lungs, skin, & mucous membranes.
- Pulmonary absorption commonest & fastest.
- Smokeless tobacco absorbed more gradually.
- Absorbed tobacco goes quickly to various body parts
- Nicotine intake from one cigarette varies widely,
   ranges from 10- 80 mg/day, or 0.4 -1.6 mg/cigarette
- After absorption, nicotine reaches brain within 7 seconds
- Psycho-active rewards of smoking occur quickly & is highly reinforced.



# MECHANISM OF ACTION



- Nicotine binds to nAChRs in brain & influences
   cerebral metabolism & increases transmitter release .
- Chronic administration leads to desensitization & inactivation of nAChRs with subsequent up-regulation of nAChRs sites.



# nAChRs & OTHER RECEPTORS

- nAChRs are concentrated in midbrain areas:
  - Mid-brain
  - > Tegmentum,
  - > Striatum,
  - Nucleus accumbens,
  - Ventral tegmental area (VTA),
  - Also muscles, adrenal glands, heart, & other organs.
- Nicotine also binds to cholinergic receptors in:
  - Autonomic ganglia, adrenal medulla,
  - Chemoreceptors of carotid bodies & aortic body & neuromuscular junction.
- Nicotine also binds to receptors in:
  - Nigrostriatal mesolimbic dopaminergic neurons.



# **NICOTINE-ACETYLCHOLINE RECEPTORS**

- Nicotine receptors have 12 subunits:
  - 9 alfa (alfa<sub>2</sub> to alfa<sub>10</sub>) &
  - > 3 beta (beta<sub>2</sub> to beta 4) have central role in autonomic transmission.
- nAChRs complex has 5 subunits in peripheral & CNS system.
- Most abundant receptor subtypes are:
  - alfa<sub>4</sub>-beta<sub>2</sub>,
  - ➤ alfa<sub>3</sub>-beta<sub>4</sub> &
  - alfa7 (homomeric).
- Beta subunit has role in nicotine addiction.
- □ Alfa<sub>4</sub>-beta<sub>2</sub> combination has greatest sensitivity to nicotine.
- Alfa<sub>3</sub>-beta<sub>4</sub> & alfa7(homomeric) mediate CVS nicotine effects.
- Alfa<sub>7</sub> also involved in rapid synaptic transmission & plays role in learning & sensory gating.



## MID-BRAIN DOPAMINE REWARD PATHWAY

- Nicotine is one of most potent stimulants of midbrain dopamine' reward pathway & when stimulated, there is release of
  - > Acetylcholin,
  - Norepinephrine,
  - Dopamine
  - Serotonin,
  - Vasopressin,
  - Growth hormone and ACTH.
- Nicotine acts on locus ceruleus regulating vigilance, arousal, concentration & stress reactions making tobacco users more alert.
- Interaction between nicotine & nAChRs affects learning, memory and other functions.



# NEUROTRANSMITTER RELEASE & BEHAVIOURAL EFFECTS

Dopamine	Pleasure & Appetite suppression	
Norepinephrine	Arousal & Appetite suppression	
Acetylcholine	Arousal & Cognitive enhancement	
Glutamate	Learning & memory enhancement	
Serotonin	Mood modulation & Appetite suppression	
Beta-endorphin	Reduction of anxiety & Tension	
GABA	Reduction of anxiety & Tension	



## NICOTINE AS AN ADDICTIVE

- Nicotine leads to tolerance to its own action like other dependence producing drugs.
- Use of 4 cigarettes is sufficient to expose a person to the risk of becoming dependent
- Within few days symptoms of dependence start appearing.
- Because of addiction tobacco consumption continues, leads to high morbidity & mortality.



# **NICOTINE WITHDRAWAL SYMPTOMS**

- Worst in first 24-48 hours gradually decline in intensity disappear in few weeks
- Symptoms may include 4 (or more) of the following within 24 hours of stopping of smoking, often causing significant distress.
  - Craving,
  - Depressed mood,
  - Increased appetite or weight gain,
  - Irritability, frustration, anger,
  - > Anxiety, difficulty in concentration, restlessness.



# TOBACCO SMOKING CESSATION

- Tobacco-related deaths are preventable.
- "Nicotine dependence warrants medical treatment as does any drug-dependence disorder or chronic disease." (Fiore et al, 2000)
- Physicians can reduce morbidity & prevent mortality.
- Every visiting patient should be asked about smoking history & possibility of leaving it.
- Methods used for tobacco-cessation include:
  - Pharmacotherapy,
  - Behaviour therapy,
  - Counseling, independently or in combination



## TOBACCO DEPENDENCE IS

# Assess the following:

- Detailed history,
- Motivation to leave
- Physical & mental health status & co-morbidity
- Environmental factors working as re-inforcers
- Benefits of cessation and possible withdrawal symptoms be explained.
- Severity of addiction



## ASSESSMENT OF NICOTINE DEPENDENCE

#### FAGERSTROM TEST FOR NICOTINE DEPENDENCE

- How soon after you wake up do you smoke your 1st cigarette? (Within 5 min.=3;5-30 min.=2;31-60 min.=1 and beyond 60 min.=0)
- Do you find it difficult not to smoke at places where you should not? (Yes=1; No=0)
- Which cigarette you would you most hate to give up?
   (First cigarette in the morning=1; Any other one=0)
- How many cigarettes do you smoke each day? (10 or fewer=0; 11-20=1; 21-30=2 and 30 and beyond=3)
- Do you smoke more during the first few hours of waking? (Yes=1; No=0)
- Do you still smoke if you are so sick that you are in bed most of the day? (Yes=1; No=0)
- SCORING: 7-10 points=High dependence; 4-6= Moderate; Less than 4= Minimal



# Time to first cigarette (TTFC)





# TTFC Less than or equal to 30 minutes after waking = HIGH DEPENDENCE



# DIFFICULTIES IN TOBACCO CESSATION

#### CUES TO SMOKING:

Some people smoke when they work alone, others smoke in company. Sight of ashtray, other person smoking, some smoke when tense, frustrated, angry, some smoke to regulate their mood.

#### BARRIERS TO SMOKING CESSATION:

- > ADDICTION:
  - Main barrier,— Irritability, restlessness, poor concentration, hunger, craving
- > FEAR OF WEIGHT GAIN:
  - Smoking increases metabolism, decreases serum leptin level, it decreases appetite and reduces body weight
- > DEPRESSION:
  - 2 times more common in women, smoking 4 times more common in depression,
     Nicotine improves mood, some smokers self-medicate to relieve depression.
- > STRESS:
  - Family environment, social relations, disadvantaged group of people.



# MEDICATION FOR SMOKING CESSATION

FIRST-LINE	DOSAGE	ADMINISTRATION
BUPROPIONE— non- nicotine, block dopamine & nor-adrenaline, & increases their level.	150 mg	150 mg daily for 3 days and then 150 mg twice a day for 7 – 12 weeks. Quit date 1-2 wks after start
NICOTINE GUM nicotine absorbed by oral mucosa, chewed, kept under tongue	2 mg piece for 25cigarettes, 4 mg piece for more than 25cigarettes per day.	1 piece every 1-2 hrs. for 6 wks, 1piece every 2-4 hrs. for 2 wks, 1 piece every 4-8 hrs for 2 wks, max. 24 piec.
TRANS-DERMAL PATCHES	21 mg/24 hours, 14 mg/24 hours and 7 mg/24 hours	>10 cigarettes/day, 21 mg for 6 wks, then 14 mg for 2 wks & 7 mg/day for 2 wks
NICOTINE LOZENGE	2mg pce if <25 cigarettes/per day, 4 mg pce if >25 are smoked.	1 pce 1-2 hrs for 6 wks, pce every 4 hrs for 2 wks & then 1 piece every 4-8 hrs
NICOTINE SPRAY	1 mg-spray each nostril	1-2 spray/hr 40 doses/day & max. 3 months use.
NICOTINE INHALER	6-16 cartridges	4 mg metered spray.



# MEDICATION FOR SMOKING CESSATION

SECOND-LINE	DOSAGE	ADMINISTRATION
adrenergic receptor agonist, anxiolytic. Can cause hypotension & bradykinesia	0.15 075 mg/day	01, 02 and 0.3 mg tablets
NORTRIPTYLINE – TCA, blocks reuptake of norepinephrine	75mg – 100 mg/day	10, 25, 50 and 75 mg tablets.



# Non-nicotine agents

#### Varenicline:

- Has high & selective activity at alfa4-beta2 receptor.
- A partial agonist at this receptor in vivo producing lesser response than that of nicotine (30-60%) but also blocks the effect of any nicotine added to the system.
- Maintains a moderate level of dopamine release, which reduces craving & withdrawal during abstinence.
- Blocks reinforcing effects of nicotine obtained from cigarette smoke in case of relapse.



# Non-nicotine agents

#### NicVAX:

- It works by stimulating immune system to make antibodies that bind to nicotine molecules, making them too big to cross the blood-brain barrier & preventing them from reaching nicotine receptors & trigger pleasure sensation that smokers & users of nicotine experience & become addicted to.
- Data from pre-clinical trials suggest that injectable vaccine is effective not only in helping people quit smoking but also from relapsing back because the nicotine antibodies last a long time.



# BEHAVIOURAL TREATMENT FOR NICOTINE DEPENDENCE

- Aversion therapy
  - Satiation
  - Rapid smoking
- Contingency contracting
- Social support
- Coping skills training
- Relaxation training



# **OTHER THERAPIES**

- Hypnotherapy: is used to weaken the desire to smoke
   & strengthen will to stop smoking through posthypnotic suggestions that:
  - Smoking is poison to your body
  - You need your body to live
  - You owe your body this respect and protection.
- Acupuncture: involves the use of needles or staple like attachments mainly at the pinna of ear.



# PHYSICIAN'S COUNSELING

#### SMOKING CESSATION: STRATEGIES, "THE 5 A's" \*

- ASK Systematically identify all tobacco smokers at every visit.
   Implement an office-wide system that allows the inquiry and documentation of tobacco use for every patient.
- ADVISE Strongly urge all smokers to quit in a strong, clear and personalized manner.
- ASK every smoker if he or she is willing to make a quit attempt at this time.
- ASSIST- Aid the patient in quitting, help patient with the development of a quit plan. Encourage nicotine replacement therapy and/or bupropion.
- ARRANGE- schedule follow-up contact in person or via telephone.



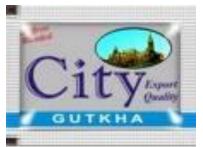
# **TOBACCO: A DEADLY SUBSTANCE**

- A <u>public health danger</u> of modern times.
- Tobacco smoke <u>kills</u> the smoker silently <u>much before</u> the actual age to exit.
- Smokers harm not only themselves, they <u>expose others</u> to the risk of morbidity and mortality.
- Smokers are unaware that they raising <u>sick and unhealthy</u>
   <u>children</u> in their houses.



# What do these colourful packets look like? Poison sachets or mouth freshners?















They are fooling the children of our country and we are allowing them to!



# 1 CIGARETTE -7 Minutes of Your Life THANKS



