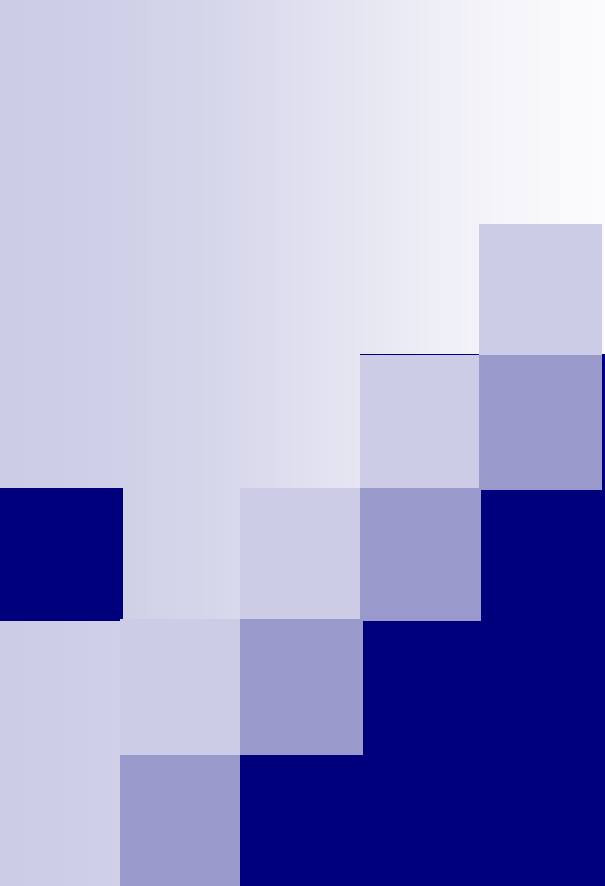


Community Medicine  
Medical Research  
Nicotine Addiction and NRT

Dr's Summary with Student own hand notes.

Written by : Mousa Al-Neimat



# Nicotine addiction and replacement therapies

Dr Munir Abu-Helalah MD MPH PhD  
Department of Family and Community Medicine  
School of Medicine, University of Jordan

Associate Professor of Epidemiology  
and Preventive Medicine  
Consultant Preventive Medicine

# Effects of smoking on your body

Smoking causes or increases your risk of:

- Macular degeneration.
- Cataracts.
- Vision loss.

- Reduced taste and smell.
- Tooth staining.
- Tooth decay.
- Gum disease.

- Frequent infections.
- COPD.

- Erectile dysfunction.
- Infertility.



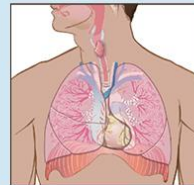
- Miscarriage.
- Birth defects.
- Developmental delays.

- Anxiety.
- Depression.
- Irritability.

- Blood clots.
- Heart attack.
- Stroke.

- Bone fractures.
- Osteoporosis.

- Premature aging.
- Yellow nails.
- Wrinkles.



- Lung, oral, laryngeal and other cancers.

# Vape and Electronic cigarettes risks:

EVALI (E-cigarette, or Vaping, Product Use-Associated Lung Injury)

- **What it is:** A serious and sometimes fatal acute or subacute respiratory illness.

- **Symptoms:** Shortness of breath, cough, fever, vomiting, diarrhea, headache, dizziness, rapid heart rate, and chest pain.

- **Cause:** While EVALI was initially strongly linked to vitamin E acetate in illicit THC-containing products, the term is used to describe a spectrum of lung injuries from vaping.

- **Diagnosis:** There is no single test; diagnosis is made by ruling out other conditions and is based on the presence of lung infiltrates on a chest X-ray or CT scan, occurring within 90 days of vaping.

THC المنتج غير القانوني المحظور في

دوخة



# Vape and Electronic cigarettes risks:

Other vaping-related lung conditions

- **Bronchiolitis obliterans:** More commonly known as "popcorn lung," this is a condition where the small airways in the lungs become inflamed and damaged.
- **Inflammation and irritation:** Vaping can cause inflammation and irritation in the lungs and airways.
- **Lung scarring:** There is evidence that vaping can lead to long-term lung scarring.

ندبات في الرئة

stiffness and hardness in lung tissue which make it hard for it to expand.



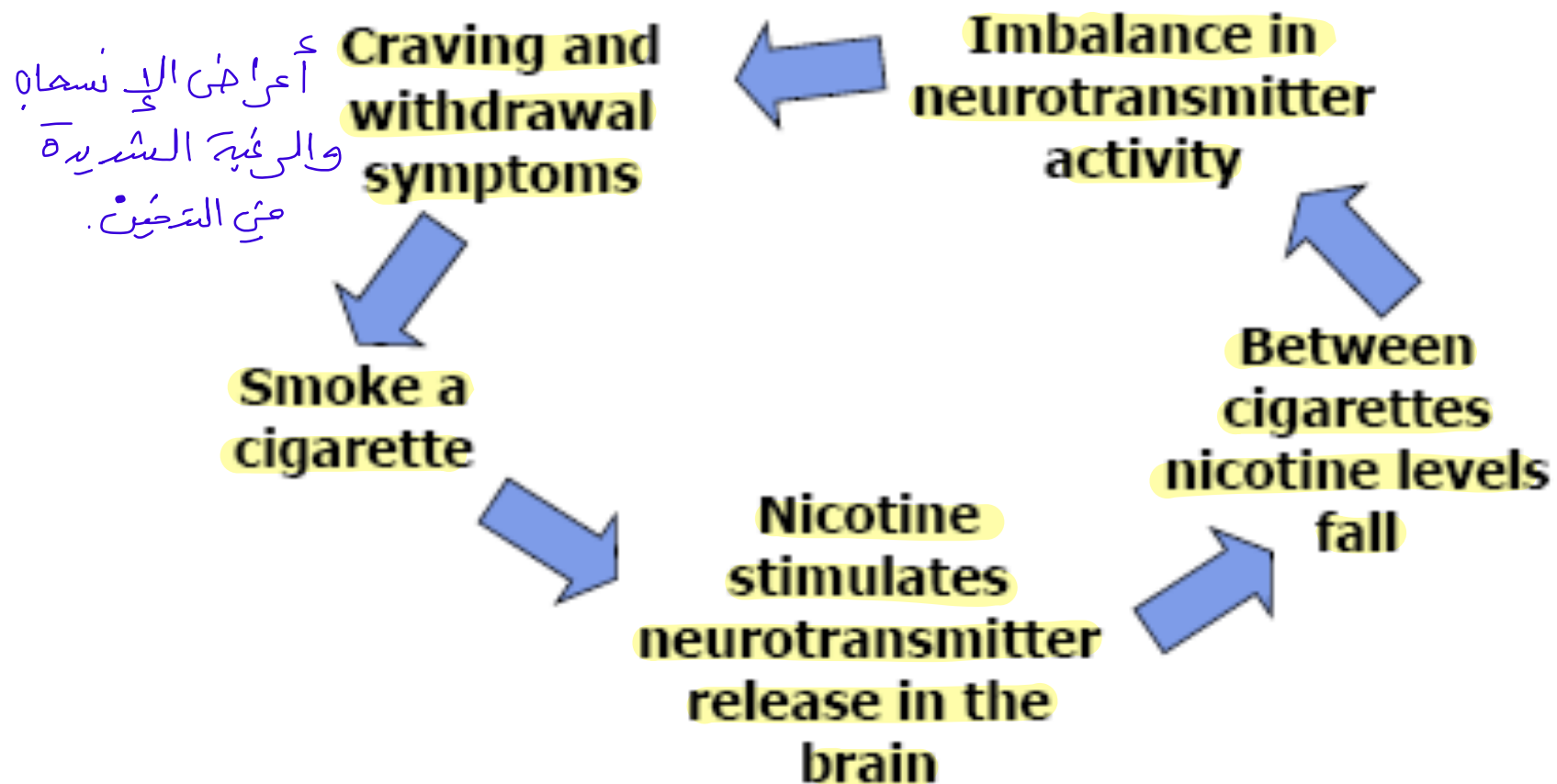
# Smoking and Brain Development

- Smoking while the brain is still developing until about age 25 can disrupt brain maturation, leading to long-term issues with attention, impulse control, learning, and memory.
- Nicotine alters brain structure and function, potentially causing permanent changes and increasing the risk of addiction to other drugs

# Addiction to nicotine

- An understanding of how nicotine produces addiction and influences smoking behavior provides a necessary basis for smoking cessation therapies.
- Chronic nicotine exposure results in neuroadaptation, that is, the development of tolerance. *Addiction*  
*Not being sensitive to the*  
*Same amount of nicotine.*
- Neuroadaptation is associated with an increased number of brain nicotinic cholinergic receptors.
- Chronic exposure to nicotine also results in changes in gene expression and neural plasticity; which is defined as “ability of the brain to reorganize neural pathways based on new experiences”

# Nicotine Addiction



# BIOLOGY of NICOTINE ADDICTION: ROLE of DOPAMINE

Take a cigarette

Nicotine  
stimulates  
dopamine release

Pleasurable feelings

Repeat administration

Tolerance develops

Nicotine addiction  
is ***not*** just a bad habit.

*Discontinuation leads to  
withdrawal symptoms.*

# Dopamine

نشوة

- **Dopamine** induces feelings of euphoria and pleasure and is responsible for activating the dopamine reward pathway
- The dopamine reward pathway, as depicted in this simplified diagram, is a network of nervous tissue in the middle of the brain that elicits feelings of pleasure in response to certain stimuli. → Dopamine

# Addiction to nicotine

*nicotine containing products,  
(it's the name of a plant).*

*مصدر العوازل*

- **Addiction to tobacco is multifactorial:**

- It includes a desire for the direct pharmacologic actions of nicotine, relief of withdrawal symptoms, and learned associations.

*يهرب من أعراض  
الانسحاب.*

- Smokers usually provide different reasons for smoking that could include pleasure, arousal, enhanced vigilance, improved performance, relief of anxiety or depression, reduced hunger, and control of body weight

*تنشيط الجهاز  
الحيوي مما يؤدي  
إلى اليقظة (Vigilance)*

*Same as  
arousal*

# NICOTINE PHARMACODYNAMICS: WITHDRAWAL EFFECTS

- Depression
- Insomnia أرق
- Irritability/frustration/anger تهيج تهيج عصب
- Anxiety
- Difficulty concentrating
- Restlessness as a result.
- Increased appetite/weight gain الشهية
- Decreased heart rate
- Cravings\* الرغبة الشديدة في العودة للتدخين.

Most symptoms peak 24–48 hr after quitting and subside within 2–4 weeks.



# What are the benefits of stopping smoking?

- It is never too late to stop smoking to gain health benefits.
- For example if the smoker already has a COPD or a heart disease, there would be great improvement in prognosis upon giving up smoking.
- Smoking cessation for smokers with history of ischaemic heart disease is expected to reduce risk of a subsequent fatal heart attack by 25 per cent.

## Timeline of health benefits after stopping smoking

Time line (After ...)	<u>Health Benefit</u>
8 hours	<u>Nicotine and CO in blood drop by 50%</u>
After 24 hours	<u>Lung starts to clear debris</u> ← معطام الخلايا وكذا.
After 48 hours	<u>Smell and taste improve</u>
72 hours	<u>Breathing becomes easier. Bronchial tubes begin to relax and energy levels increase.</u>
1 month	<u>Skin appearance improves</u> , <sup>لبيسب</sup> <u>owing to improved skin perfusion.</u>
3-9 months	<u>Cough, wheezing, and breathing problems improve and lung function increases by up to 10%.</u>
1 year	<u>Risk of a heart attack falls to about half that of a smoker.</u>
10 years	<u>Risk of lung cancer falls to about half that of a smoker.</u>
15 years	<u>Risk of heart attack falls to the same level as someone who has never smoked.</u>

<p>Nicotine &amp; CO ↓ by 50% in Blood</p> <p>8</p> <p>24</p> <p>Lung start to clear debris</p>	<p>Tasting &amp; smelling improves</p> <p>48</p>	<p>Skin appearance improves due to improve skin perfusion.</p> <p>72</p> <p>Breathing become easier and bronchial start to relax and increase in the level of energy.</p>	<p>1</p>	<p>3-9</p> <p>Lung function improved by 10%</p> <p>cough, wheezing &amp; Breathing problems improves.</p>	<p>1</p>	<p>10</p> <p>lung cancer risk ↓ to a level that is half to smoker</p>	<p>15</p> <p>Heart attack risk ↓ level of person who had never smoked</p>
---	--	---	----------	---	----------	---	---

## Modified Fagerstorm Scoring:

*Most  
important*

1. How soon after you wake up do you smoke your first cigarette?
  - Within 5 minutes (3 points)
  - 5 to 30 minutes (2 points)
  - 31 to 60 minutes (1 point)
  - After 60 minutes (0 points)
2. Do you find it difficult not to smoke in places where you shouldn't, such as in church or school, in a movie, at the library, on a bus, in court or in a hospital?
  - Yes (1 point)
  - No (0 points)
3. Which cigarette would you most hate to give up; which cigarette do you treasure the most?
  - The first one in the morning (1 point)
  - Any other one (0 points)
4. 4. How many cigarettes do you smoke each day?
  - 10 or fewer (0 points)
  - 11 to 20 (1 point)
  - 21 to 30 (2 points)
  - 31 or more (3 points)
5. 5. Do you smoke more during the first few hours after waking up than during the rest of the day?
  - Yes (1 point)
  - No (0 points)
6. 6. Do you still smoke if you are so sick that you are in bed most of the day or if you have a cold or the flu and have trouble breathing?
  - Yes (1 point)
  - No (0 points)

**Scoring:** 7–10 points = highly dependent; 4–6 points = moderately dependent; less than 4 points = minimally dependent.

# Pack-Year

- A pack-year is a clinical quantification of cigarette smoking used to measure a person's exposure to tobacco.
- Used for risk assessment
- Definition
- The pack-year is a unit for measuring the amount a person has smoked over a long period of time. It is calculated by multiplying the number of packs of cigarettes smoked per day by the number of years the person has smoked.
- For example, 1 pack-year is equal to smoking 20 cigarettes (1 pack) per day for 1 year
- One pack-year is the equivalent of 365 packs of cigarettes or 7,300 cigarettes, in a year as smoker.

# Pack-Year

## Calculation

- Number of pack-years = (packs smoked per day)  $\times$  (years as a smoker)
- or
- Number of pack-years = (number of cigarettes smoked per day/20)  $\times$  number of years smoked. (1 pack has 20 cigarettes in some countries)
- Note that despite the unit being called a "pack-year," the actual unit is simply a number of packs (as noted above).
- 1 pack-year = 1 pack/day  $\cdot$  1 year = 1 pack/day  $\cdot$  365 days = 365 packs = 365 packs  $\cdot$  20 cigarettes/pack = 7,300 cigarettes

$$\begin{aligned} 1 \text{ pack-year} &= 1 \text{ pack/day} \cdot \text{year} = 365 \text{ pack/day} \cdot \cancel{\text{day}} = 365 \text{ packs} \\ &= 365 \times 20 \text{ cigarettes} \\ &= 7300 \text{ cigarettes.} \end{aligned}$$

# Management plan:

- In the first visit:
- Medical and drug history, smoking pattern and history, nicotine dependence using Fagerstrom Tolerance Questionnaire.
- Counselling on smoking: Motivational interviewing strategies and the “5R’s” for enhancing motivation approaches will be used.
- The first approach is based on the following components: express empathy, develop discrepancy, roll with resistance, support self-efficacy. While the “5R’s” for enhancing motivation includes: Relevance, Risks, Rewards, Roadblocks, Repetition.
- Smokers, who are willing to receive smoking cessation medical therapy, would receive further assessment in order to determine the treatment of choice for their particular condition and to exclude any contraindications for the selected treatments.

## In the first visit:

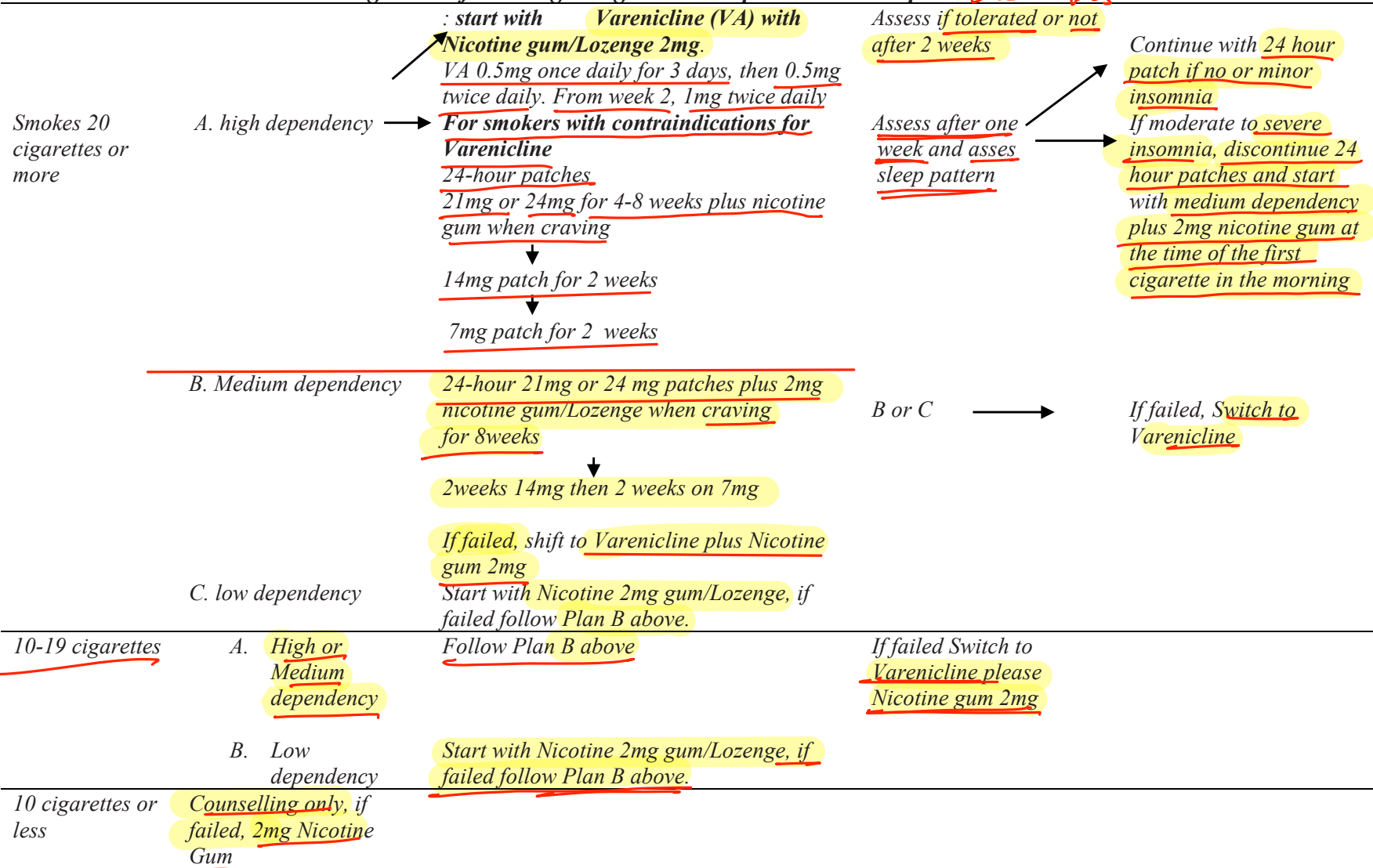
- A detailed medical and drug history will be obtained, along with an assessment of the smoking pattern and history and nicotine dependence using the Fagerström Tolerance Questionnaire.  
*Smoking Kind?* *How Long?*
- Counseling on smoking will be provided using motivational interviewing strategies and the "5 R's" approach to enhance motivation.
- The motivational interviewing approach includes the following components:
  - *Express empathy*
  - *Develop discrepancy*
  - *Roll with resistance*
  - *Support self-efficacy*
- The "5 R's" for enhancing motivation include:
  - Relevance *Personal benefits*
  - Risks
  - Rewards *Healthy benefits*
  - Roadblocks
  - Repetition *Repeat in every visit-*
- Smokers who are willing to receive medical therapy for smoking cessation will undergo further assessment to determine the most appropriate treatment for their condition and to exclude any contraindications to the selected therapy.



## في الزيارة الأولى:

- يتم جمع التاريخ المرضي والدوائي بشكل شامل، بالإضافة إلى تقييم نمط التدخين وتاريخه، وقياس مدى الاعتماد على النيكوتين باستخدام استبيان فاغريستروم لتحمل النيكوتين (Fagerström Tolerance Questionnaire).
- تُقدّم للمريض جلسة إرشادية حول التدخين تعتمد على أساليب المقابلة التحفيزية واستراتيجية "الخمس راءات (5R's)" لتعزيز الرغبة في الإقلاع.
- تتضمن المقابلة التحفيزية العناصر الآتية:
  - إظهار التعاطف مع المريض
  - توضيح التناقض بين السلوك الحالي (التدخين) والأهداف الشخصية المستقبلية
  - التعامل مع المقاومة بطريقة إيجابية دون مواجهة
  - تعزيز الثقة بالنفس والقدرة على التغيير
- أما استراتيجية الخمس راءات (5R's) فتركّز على:
  - الأهمية (Relevance): ربط الإقلاع عن التدخين بحياة المريض وظروفه الشخصية
  - المخاطر (Risks): توضيح الأضرار الصحية والاجتماعية لاستمرار التدخين
  - المكافآت (Rewards): إبراز الفوائد الصحية والنفسية بعد الإقلاع
  - العوائق (Roadblocks): مناقشة الصعوبات المحتملة ومحاولة إيجاد حلول لها
  - التكرار (Repetition): تكرار التوعية والتحفيز في كل زيارة
- أما المدخنون الراغبون في تلقي العلاج الدوائي للإقلاع عن التدخين، فيُجرى لهم تقييم إضافي لتحديد أنسب نوع من العلاج بناءً على حالتهم الصحية، والتأكد من عدم وجود موانع لاستخدام العلاج المختار.

## Guideline: management of smoking using nicotine replacement therapies إذنيها متعملة ولا





# Nicotine replacement therapy (NRT)

- Provides smoker with nicotine without using tobacco, thereby relieving nicotine withdrawal symptoms.
- NRT products differ in the route of delivering nicotine to the circulation.
- Nicotine is absorbed transdermally with the nicotine skin patch, through the nasal mucosa by the nasal spray or through the oral mucosa with the nicotine chewing gum, nicotine lozenge, or nicotine inhaler.  
*pass through dermis layer*
- Nicotine patches are more tolerated than nasal spray and provides the most continuous delivery of nicotine of nicotine replacement therapies.  
*fewer side effect (أقل أعراض) an example*

*Nicotine patches provide Nicotine continuously since it is 24 hour attached to the skin.*

*Nasal spray is a less continuous nicotine supply (أنا تبخيل فيه ينشف أو يروح) بسرعة أو غير مستمرة*

# Nicotine replacement therapy (NRT)

- Principle:

Many of the difficulties in smoking cessation stems from problems posed by nicotine withdrawal.

# Nicotine Replacement Therapy (NRT)

*Reduce*

- Reliably attenuates severity of withdrawal, making it easier for would-be ex-smokers to cope with abstinence while unlearning the deeply ingrained habit elements of smoking

*Significantly decrease the severity of withdrawal symptoms which make it easier to cope with abstinence, and we are unlearning the smoking habits itself.*

# Nicotine Replacement Therapy

- Indicated for all smokers trying to quit, except in the presence of special circumstances (those with medical contraindications, pregnant and adolescent smokers)

المخيف المراهقين

→ Cause there is another way of leaving without NRT.

## NRT - Therapeutic Effect

- The primary therapeutic effect of NRT is to reduce the severity of symptoms associated with smoking cessation. *with drawl symptoms*
- NRT provides steady levels of nicotine and may reduce the pleasurable effects of tobacco desired by smokers. *taking needed nicotine amount with reduced pleasure and reduce nicotine concentration over time*
- NRT makes it easier to cope in difficult situations.



# Nicotine patches

- Transdermal patch (waterproof)
- Nicotine Replacement Therapy (NRT)
- To overcome withdrawal symptoms
- 24-hour patch to deliver constant nicotine levels
- *8 weeks with 21mg → 14mg for 2 weeks → 7mg for 2 weeks*  
12 week weaning program that reduces and eliminates body's dependence for nicotine
- Clinically-proven to be better than *قوة الدخان* willpower alone
- 2-4 times the success rate of placebo patches

*لله مقالة وحقيرة*

Hajek P, West R, Foulds J, Nilsson F, Burrows S, Meadow A. Randomized comparative trial of nicotine polacrilex, a transdermal patch, nasal spray, and an inhaler. *Arch Intern Med.* 1999;159:2033-2038.

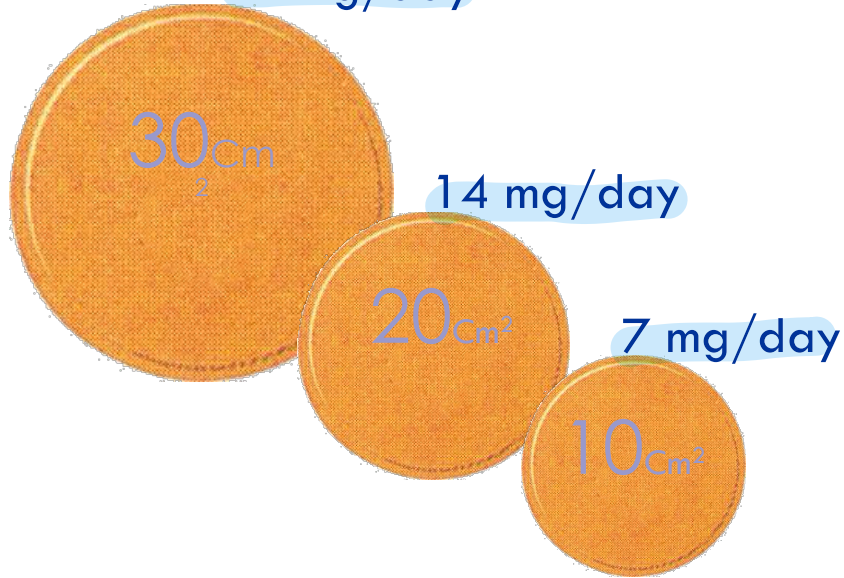
# Nicotine patches

for then for then for  
21 or 24mg/24hrs, 14mg/24hrs, 7mg/24hrs

for  
21 or 24mg/16hrs, 14mg/16hrs, 7mg/16hrs

the rest of the day without patches  
15mg/24hrs, 10mg/24hrs, 5mg/24hrs

21 mg/day



*Different courses of  
nicotine patches.*

# Precautions

- ❑ women who are pregnant or breast feeding
- ❑ smokers with cardiovascular conditions

Doctors should weigh risks/ vs benefits when prescribing NRT to pregnant women or smokers with cardiovascular conditions.

- ❑ smokers using other nicotine replacement products

*As a doctor this should be kept in mind.*

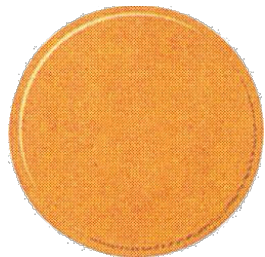
*of it  
could have severe  
side effect*

- ❑ children
- ❑ non-smokers
- ❑ smokers of fewer than 10 cigarettes a day

*start with consulting.*

# Nicotine patches– Application

- ❑ Apply to non-hairy, clean, dry skin
- Rotate between sites (this helps to reduce the risk of skin irritation) *تدوير بين المواقع*
- Choose a flat surface *مسطح توضع عليها*
- Avoid joints or skin folds
- Replace the same time everyday



**DO NOT SMOKE WHILE  
USING THE PATCH**

# Why a 24-hour patch ?

- ▶ Many of the dependent smokers
  - Either smoke just before to go to bed,
  - Or wake up at night for smoking
  - Or wake up early in the morning to have a cigarette

Night smoking is a criteria for strong addiction to  
nicotine

People who wake up at night  
or take a cigarette before  
going to bed.

# Nicotine Lozenges

- Nicotine Lozenges are new products that have similar efficacy to nicotine gum but it is easier to use and does not require special technique for optimal use.
- It also delivers more nicotine than equivalent dose of the gum.
- Unlike nicotine gum, smokers with dentures or poor dentition can use Lozenges.

اسنان متحركة او خرابة .

ضعف الاسنان


# High level of smoking addiction

Combination of varenicline plus Nicotine gum with behavioural therapy

If Varenicloine is contraindicated:

Combination of behavioural therapy with nicotine patches and nicotine lozenges (used for cravings)

لما تولع حطك وبك تدخن.



**For patients with ischaemic heart disease with moderate or high level of addiction:**

The best approach is to start with **behavioural therapy**

If failed, you can start with nicotine patches



# Varenicline (Chantix, Chantix)

- a partial agonist at the  $\alpha 4 \beta 2$  subunit of the nicotinic acetylcholine receptor.
- This drug binds to nicotinic receptors leading to reduction of withdrawal symptoms and decrease rate of cigarettes' nicotine binding to these receptors.

# Varenicline (Chantix, Chantix)

*the absence of nicotine withdrawal symptoms*

- This medication has shown a good abstinence rate when compared to nicotine patches, but the difference is small when compared to combined nicotine patches with gum or lozenges.
- The best scientific evidence supports combination of Varenicline with Nicotine gum.
- Combining varenicline and nicotine gum is associated with higher rates of continuous abstinence at 12 weeks (55.4% vs. 40.9%) and 24 weeks (49.0% vs. 32.6%) compared to varenicline alone.

# Varenicline (Champix, Chantix)

- Week 1: Starting dose: 0.5mg once daily for three days, then 0.5mg twice daily for four days.

Smokers have to decide on quit day during this week.

- Then  
1mg twice daily for 8-12 weeks.

# Contraindications

*You should not prescribe Varenicline if smoker has one of these.*

- Varenicline has not been studied in children and should not be taken by young people who are under 18 years of age.
- Breast feeding. Varenicline may pass into breast milk.

Either use an alternative therapy or follow other ways of feeding the baby may be appropriate if she is currently taking the drug.

## Precautions for varenicline use

*its better to not prescribing vareniclin.*

- kidney problems or on dialysis.
- Pregnancy. The effects of Varenicline on the foetus are not known and it would be better to initiate an alternative therapy if behavioural intervention is not successful.



## Precautions for varenicline use

- Depression or any psychiatric illnesses

# Varenicline: Side effects

- Vomiting and nausea داسفرائی غیاں ناوسیا
- Headaches
- Sleep disturbances and atypical dreams
- Gas (wind)
- Changes in the way food tastes  
(Dysgeusia) دس جیوٹسا
- Constipation
- Suicidal thoughts

# Bupropion (Zyban)

- Available as an antidepressant in the United States since 1989, is believed to act by enhancing central nervous system noradrenergic and dopaminergic release.
- A sustained-release formulation of the drug is licensed as an aid to smoking cessation (Zyban); it is identical to the antidepressant Wellbutrin SR and is available as a generic drug.





# Bupropion (Zyban)

- A meta-analysis of 31 randomized trials of bupropion monotherapy concluded that bupropion SR doubles the likelihood of smoking cessation

# Bupropion (Zyban)

- As an example, one multicenter, randomized, double blind trial of 615 patients compared sustained-release bupropion (150 mg twice daily) with placebo among patients who wished to stop smoking.
- The rates of smoking cessation (confirmed by exhaled carbon monoxide measurements) were significantly greater at the end of a seven-week course of treatment among patients who received bupropion (44 versus 19 percent).

# Bupropion (Zyban)

- **Safety** — The most common side effects of bupropion are insomnia, agitation, dry mouth, and headache. مشکل جی (بیمور ہو شات)
- A more serious side effect is seizure, which can occur because bupropion reduces the seizure threshold. نوبات
- In clinical trials, the risk of seizure was 0.1 percent, and the drug is contraindicated in patients with a seizure disorder or predisposition to seizure. محرکین للنوبات

إدعوا لنا بظهر الغيب...