#### Acne : An Update

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### Conflict of interest

- Advisory board :
- Galderma
- Johnson and Johnson
- Pfizer
- Byer
- Sunpharma



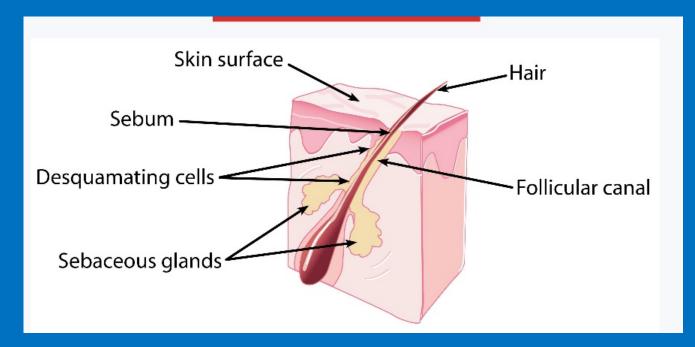
- As a result of attending this session, participants will be able to:
- 1-Understand the pathophysiology of Acne vulgaris in different age groups
- 2-Prescribe appropriate therapy according to the patient's clinical presentation
- 3-Recognize the role of diet in acne

# Epidemiology

- Acne is the most common skin disorder
- Affects 85% of young people between 12 and 24 years of age
- Typically presents at ages 8-12 Y, peaks at ages 15-18 Y
- May resolve by 20-25 years of age
- May persist till 50 years of age

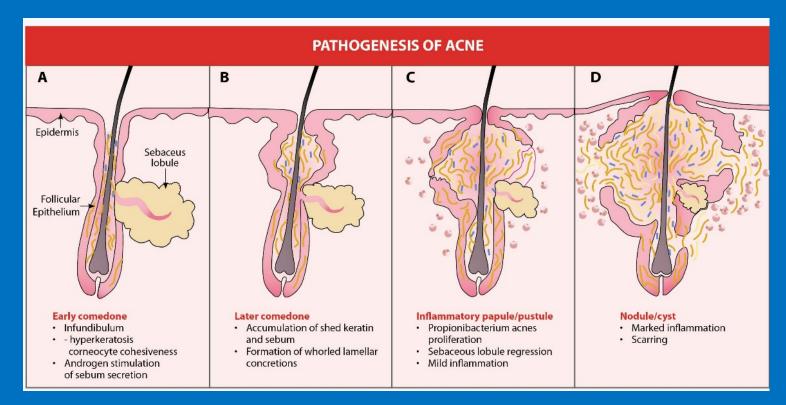
### Acne

- Chronic, inflammatory disease of the pilosebaceous units
- Pilosebaceous unit= hair + hair follicle + sebaceous gland



### Acne

- Excessive sebum production
- Abnormal follicular keratinization
- Proliferation of Proprionibacterium acnes
- Inflammation



## How to diagnose acne?

- By identification of lesions
- Comedones
- Papule
- Pustule
- Nodule

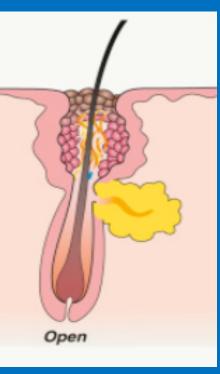
### **Open Comedone**

• Slightly elevated papules with a dilated follicular outlet filled with blackened keratin

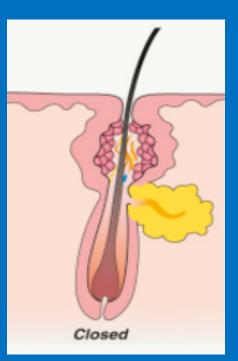
#### **Closed comedone**

• Small skin-colored papules with no apparent follicular opening









### Trichostasis spinulosa

• Clusters of vellus hairs become embedded within hair follicles, with resultant dark, spiny papules on the face or trunk.



IDOJ.2014,5 (6):132-13

#### Papule

• Solid raised lesion less than 1 cm



#### **Pustule**

• Elevated lesion containing pus



#### Nodule

 Palpable solid lesion larger than 1 cm



### Why to treat?

Acne often spontaneously clears

Psychological impact :

Emotional distress, decreased self-esteem, depression

Permanent scarring







https://acne-scar-treatment.com/scar-revision

### Post inflammatory color changes





- See the acne in context
- Is this normal for the patient age group to have acne?

### Neonatal acne

- May be present at birth, or develop over first 6 weeks of life
- Not a real acne but Inflammatory reaction to Malassezia furfur/ no comedone
- Inflammatory erythematous papules and pustules on face and scalp
- Resolves spontaneously and without scar within the first 3 months of life
- May respond more quickly with topical anti-yeast (e.g. ketoconazole cream)



### Infantile acne

- Infantile acne initially presents at 2–12 months of age
- True acne with comedones



- Spontaneous resolution in 6-12 months of age
- Elevated LH in first year of life in boys and enlarged fetal adrenal gland in both sexes lead to increased androgens
- Hormonal work up if sever, persistent or associated with other signs of hyperandrogenism

### Childhood acne

- Acne presenting between 1 and 7 years of age
- MOST WORRISOME!
- No normal source of androgen at this age group



• Think of : Premature adrenarche, Cushing's syndrome, CAH, gonadal/adrenal tumors, precocious puberty

#### Pre-adolescent acne



- Acne presenting between 7 and 12 years of age
- Androgen produced by adrenal glands as part adrenarche
- May precede other signs of pubertal maturation
- Mostly comedonal acne of forehead and midface
- Work-up beyond history and physical is generally unnecessary unless other signs of androgen excess, polycystic ovarian syndrome, or other systemic abnormalities

## Adolescent acne

- 90% of adolescents develop acne to some extent
- Androgen hormones production during puberty
- In females adolescents with sever acne always consider the PCOS

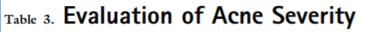
#### Post- adolescent/adult acne

- Acne that occurs after age 25
- It can be persistent, with onset during teenaged years, or late onset beginning after the age of 25 years
- Smoking, cosmetics, stress and sleep deprivation may contribute
- Red flags:
- Hair loss
- Excess hair growth
- Irregular menstrual cycles
- Rapid weight gain or loss
- Rapid onset of acne with no prior history of acne



#### Management

- Now determine the type and severity of acne
- Comedonal vs inflammatory acne
- Scarring vs nonscarring



| Grading  | Predominant Lesion Type  | Distribution                              | Scarring   | Other Factors                        |
|----------|--|---|--|--------------------------------------|
| Mild     | Few to several comedones, few scattered<br>papules   | <1/4 <sup>th</sup> face, mostly<br>T zone | None   | None                                 |
| Moderate | Many papules and pustules, variable<br>comedones, 1-2 nodules                                  | Roughly 1/2 face                          | Few, shallow   | Involvement of the<br>chest and back |
| Severe   | Numerous papules and pustules and<br>nodules; variable comedones; sinus<br>tracts and/or cysts | Face, back, and/or<br>chest               | Moderate to<br>extensive,<br>hypertrophic<br>and/or deep | Drainage, hemorrhage,<br>pain        |

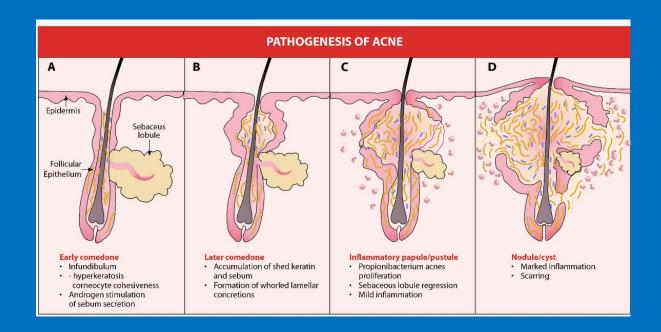






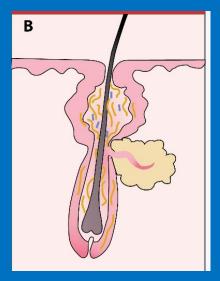
### Acne therapy

- Anti-seborrheic
- Keratolytic/comedolytic
- Antibacterial
- Anti-inflammatory



# **Topical Retinoids**

- Vitamin A derivatives
- Normalization of follicular keratinization and anti-inflammatory
- Considered as first line of TX for acne
- Monotherapy for comedonal acne
- Combinational therapy for inflammatory acne



### Natural Topical Retinoids

- Tretinoin (retinoic acid ) the acid form of vitamin A
- 0.01-0.1 concentration
- SE: Dryness, scaling, erythema, burning, irritation, and photosensitivity
- Apply retinoid every other day for first 2-4 weeks or use a short contact method for first 2-4 weeks
- Can be degraded if used during the day or in conjunction with benzoyl peroxide

### Tretinoin

- Stieva-A
- Retin A
- Retin A micro gel microsponge release formulation (0.1 and 0.04% Gel)









## Synthetic Retinoid

- Adapalene (Differin)
- Less irritant
- Seems more effective





### Synthetic Retinoid

- Tazorac (Tazaroten)
- Cream and gel 0.0,5 and 0.1



### Synthetic Retinoid

• AKLIEF (Trifarotene)



### Retinoids

| Stieva A                    | 45gr  | 25 \$* DE 20 |
|-----------------------------|-------|--------------|
| Retin A                     | 30 gr | 20 \$ *DE 20 |
| Retin A micro 0.04/0.1%     | 45gr  | 85\$         |
| Differin Gel /Cream<br>0.1% | 60gr  | 170 \$       |
| Differin gel XP 0.3%        | 60gr  | 175 \$       |
| Tazorac                     | 30 gr | 64 \$        |
| Aklief                      | 75 gr | 206 \$       |

# Benzoyl peroxide

- A lipophilic non-antibiotic antibacterial agent
- Comedolytic
- Anti inflammatory
- Comedonal and inflammatory acne
- Bleaches the clothes and hair
- Dryness and irritation

# Benzoyl Peroxide

| Benzagel                                     | 30 gr  | 10\$   |
|--|--------|--------|
| Clean and Clear Benzoyl<br>Peroxide cleanser | 150 ml | 12 \$  |
| Persagel                                     | 28 gr  | 12\$   |
| Benzac wash 10%                              | 225 gr | 100 \$ |

### Tactupump /Tactupump forte

- Adapalene(0.1%) and benzoyl peroxide topical gel(/2.5%)
- Adapalene(0.3%) and benzoyl peroxide topical gel (2.5%)



# **Topical Antibiotics**

- Antibacterial
- Anti-inflammatory
- Clindamycin
- Erythromycin
- SE: Development of resistance by P. acnes
- Resistant *P.acnes* can spread to the skin of untreated contacts
- No monotherapy
- Combination with BPO and retinoids decreases the risk of resistance



#### **Clindoxyl gel**

# • Clindamycin 1% and benzoyl peroxide 5%



#### **Benzaclin gel**

• Clindamycin 1% and benzoyl peroxide 5%



### Benzamycin

#### • Erythromycin 3% and benzoyl peroxide 5% alcohol based gel



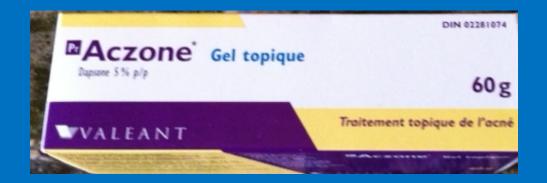


• Clindamycin 1.2% Tretinoin 0.025% aqueous gel



## Aczone gel

- Topical 5% dapsone water based gel
- Antimicrobial and anti-inflammatory effects
- Mostly for inflammatory acne



### Finacea : Azaleic acid

- A naturally occurring dicarboxylic acid produced by Malassezia furfur
- Antibacterial but not antibiotic



- Comedolytic
- It may help to lighten post inflammatory hyperpigmentation

## Salicylic acid

- A beta hydroxy-acid (BHA) derived from willow bark
- Comedolytic and antibacterial agent
- Salicylic acid is available over the counter in different products Like Reversa, Clean and Clear

| Clindoxyl gel  | 45gr  | 60\$    |  |
|----------------|-------|---------|--|
| Finacea        | 50 gr | 45\$    |  |
| Aczone         | 60gr  | 90\$    |  |
| Tactuo forte   | 60 gr | 225 \$  |  |
| Benzaclin gel  | 50gr  | 70 \$   |  |
| Benzamycin gel | 46gr  | 78\$    |  |
| Tactuo gel     | 60gr  | 140\$   |  |
| Biacna         | 60gr  | 90 \$   |  |
| Salycilic Acid |       | 10-30\$ |  |
| Dalacin T      | 60gr  | 37 \$*  |  |

## **Oral antibiotics**

- When inflammatory acne is not responding well to topical treatments and acne involving trunk or multiple bodily areas
- Should be used at least for 6-8 weeks and max up to 4 months pulse therapy
- Antibiotic resistance in P. Acne ,staph aureus and other non target bacteria
- Should be prescribed with topical therapy : Benzoyl Peroxide and/or retinoids
- Once improvement achieved scale back to topical agents alone for maintenance
- A topical retinoid and BPO or azelaic acid can be used at discontinuation of antibiotic

## **Oral antibiotics**

- Doxycycline
- SE: Dyspepsia, nausea, emesis, diarrhea, photosensitivity, esophagitis
- Dose:50-100mg once to twice daily
- Minocycline
- SE: vertigo, mouth & shin hyperpigmentation, hepatitis, drug induced lupus
- Dose: 50-100mg once to twice daily

## Hormonal therapy

Second-line treatment for female patients irrespective of whether or not the serum androgen levels are abnormal

•OCP can help with acne :

•Estrogen: Inhibit LH and FSH

Suppression of ovarian androgen production

■↑ Sex hormone binding globulin (SHBG) →  $\downarrow$  free testosterone

Not to be used until one year after onset of menstruation.

#### Oral contraceptive pills

#### All OCPs improve acne

#### Some are specifically studied

| COMMONLY USED COMBINED ORAL CONTRACEPTIVES   |  |  |  |  |
|--|--|--|--|--|
| Oral contraceptive®  | Estrogen mcg/Progestin mcg                                   |  |  |  |
| FDA-approved for acne vulgaris   |  |  |  |  |
| Ortho Tri-Cyclen   | Ethinyl estradiol 35/norgestimate 180, 215, 250              |  |  |  |
| Estrostep  | Ethinyl estradiol 20, 30,<br>35/norethindrone 1000           |  |  |  |
| Yaz, Loryna, Nikki, Beyaz*   | Ethinyl estradiol 20/drospirenone 3000                       |  |  |  |
| Clinical data to support use   |  |  |  |  |
| Alesse   | Ethinyl estradiol 20/levonorgestrel 100                      |  |  |  |
| Diane-35 <sup>†</sup>  | Ethinyl estradiol 35/cyproterone acetate 2000                |  |  |  |
| Yasmin, Syeda, Yaela,<br>Safyral*  | Ethinyl estradiol 30/drospirenone 3000                       |  |  |  |
| Natazia  | Estradiol valerate 1000, 2000, 3000/<br>dienogest 2000, 3000 |  |  |  |
| No/insufficient clinical data  |  |  |  |  |
| Various combinations of ethinyl e<br>norethindrone 400, 500, 750, 800<br>levonorgestrel 50, 75, 125, or 150<br>desogestrel 100, 125, or 150 <i>OR</i><br>norgestrel 300 <i>OR</i><br>ethynodiol diacetate 1000 | 0 OR   |  |  |  |
| *Also contains levomefolate calcium for protection against neural tube defects   |  |  |  |  |

\*Also contains levomefolate calcium for protection against neural tube defects. \*Not available in the US.

## Isotretinoin

Synthetically derived from vitamin A

Comedolytic

Decreases follicle and sebaecous gland size

Decreases sebum production

Possesses anti-inflammatory effects

### Isotretinoin

- Nodulocystic acne / Recalcitrant acne
- Total cumulative dose of 120-150mg/kg
- Common side effects : Dry mouth, nose, and eyes...
- Increases in cholesterol, triglycerides and transaminases
- Teratogenicity, Pseudotumor cerebri
- Depression ?



- Talk about temporary worsening of acne
- Mention irritation and suggest decreasing frequency of application if needed
- Don't forget moisturizers
- Gentle cleanser
- Simplify the treatment



- Realistic expectations and the importance of maintenance regimens
- Long length of treatment (Acne can last 5-10 years)
- Ensure that makeup, sunscreen, and moisturizer are non-comedogenic

| Acne Tre        | eatmei | nt Action Plan  | Hôpital de Montréal<br>pour enfants<br>Centre universitaire<br>de santé McGill | Montreal Children's<br>Hospital<br>McGill University<br>Health Centre |  |
|-----------------|--------|---|--|---|--|
|                 |        |   |  |   |  |
|                 | In the | morning,  |  |   |  |
|                 |        | Wash face with  |  |   |  |
|                 |        | Applyto face, chest, back every morning or alternating    |  |   |  |
|                 |        | Moisturize face with                                      |  |   |  |
|                 |        | Sunscreen and/or oil-free makeup can be applied after the | se steps if require  | d or desired.   |  |
| In the evening, |        |   |  |   |  |
|                 |        | Wash face with  |  |   |  |
|                 |        | Applyto face, chest, bac                                  | ck every night   |   |  |
|                 |        | Apply to face, chest, bac                                 | ck on odd nights a   | ind   |  |

## Washing and scrubbing

• Improves the oily appearance of face but does not necessarily open or cleanse pores and may lead to skin irritation

• No good study available

• Washing 2 times per day with a gentle cleanser

A single-blinded, randomized, controlled clinical trial evaluating the effect of

face washing on acne vulgaris. Pediatric dermatol 2006 Sep-Oct;23(5):421-7.

## Mask Acne : Maskne

- Mechanical acne : friction and heat, bacteria and yeast overgrowth
- What to do?
- Cotton mask
- Change masks every few hours
- Don't use make up under the mask
- Wash your face after removing mask and put non comedogenic moisturizer



# Acne and diet?

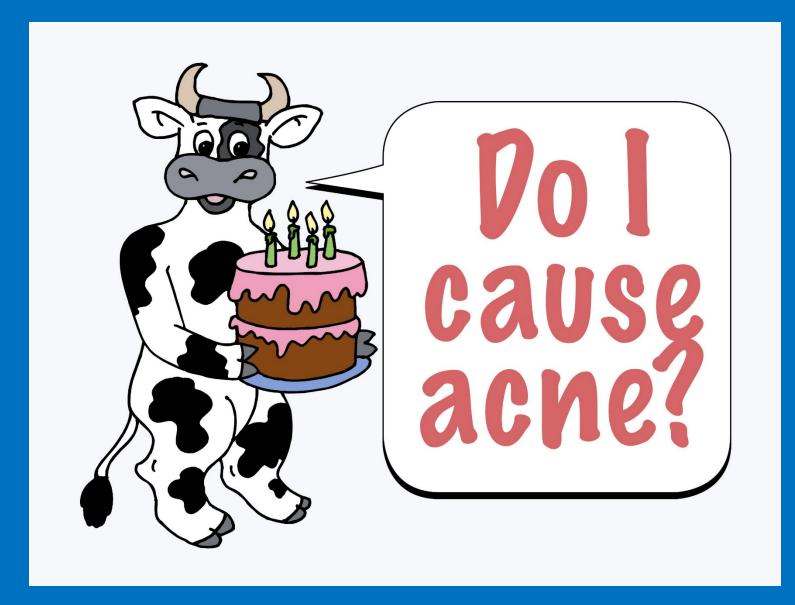




## Acne and sugar

- Diets with high glycemic load →hyperinsulinemia and increased insulin-like growth factor 1 (IGF-1) activity → androgen synthesis by various tissues as well as direct effects on pilosebaceous unit →increased sebum production and follicular plugging → worsen acne
- Multiple RCTs have shown the benefit of a low sugar diet in treating acne.
- This diet may be recommended to patients.





#### Acne and milk

- Observational studies support the link between milk and acne.
- Milk increases insulin and IGF-1 levels
- Milk contains bovine IGF-1, and dihydrotestosterone precursors
- More probably low-fat/skim milk, but not full-fat milk, is positively associated with acne
- Fat-reducing process could enhance the insulin and IGF-1-promoting elements of milk
- Milk restriction?

## Comedonal acne

- 1-Topical retinoid
- a) StiveA
- b) RetinA
- c) Differin Xp
- d) Benzagel
- d) Tactuopump



## Inflammatory acne

- Retinoid and/or benzoyl peroxide Plus/minus Topical antibiotics
- All mentioned in previous slide

Plus

- a) Benzaclin/Clindoxyl
- b) Dalacin
- c) Biacna
- If sever can add oral antibiotic



## To Take Home Messages

- Acne is the most common skin diseases encountered by primary care physicians
- Can affect mental health
- Can be scarring
- Avoid antibiotics as much as you can
- In case you need them avoid monotherapy
- Oral antibiotic for a maximum of 4 months
- Childhood acne is a real red flag
- Low glycemic diet should be emphasized

### References

- 1-Pediatrics in Review November 2013, 34 (11) 479-497
- 2-2003 Elsevier-Bolognia,Jorizzo and Rapini:Dermatologywww.dermtext.com
- 3-Am Fam Physician. 2012 Oct 15;86(8):734-740

