# **Alopecia: A Fresh Look and a New Algorithm for Diagnosis**

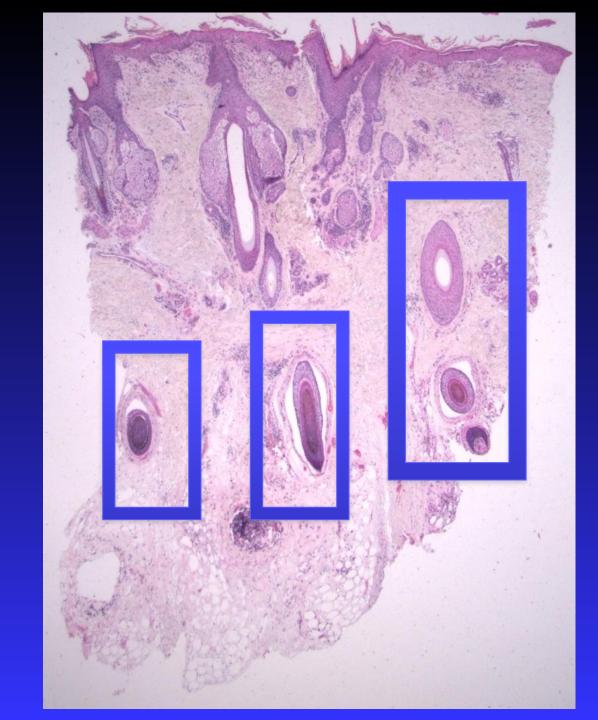
Curtis T. Thompson, M.D. Clinical Professor

Departments of Dermatology, Pathology and Biomedical Engineering Oregon Health Sciences University Portland, Oregon, USA

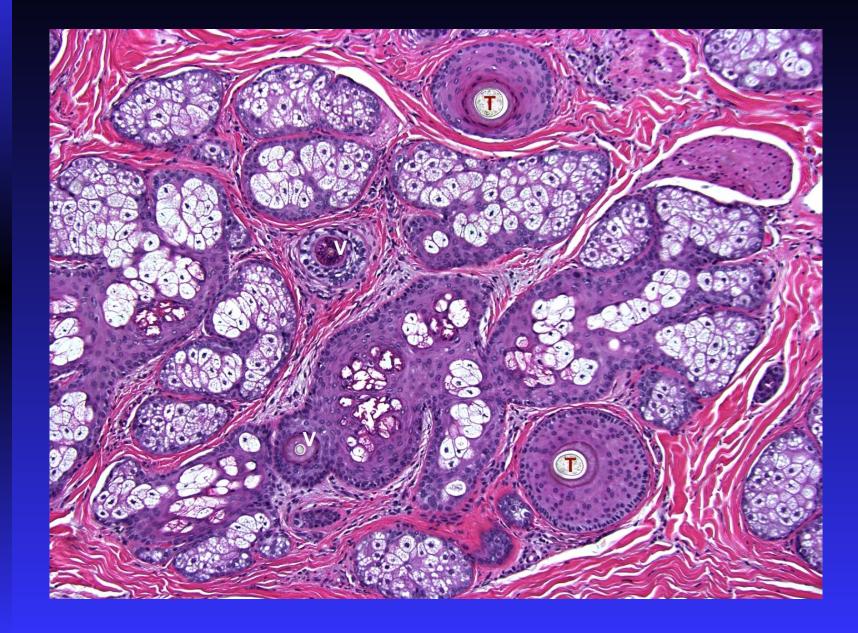
# 'Fresh' Objectives

- Tissue processing providing a more thorough examination
- H&E points of interest
  - Immunohistochemistry as an aid in diagnostic impasses
- An algorithm for better diagnostics

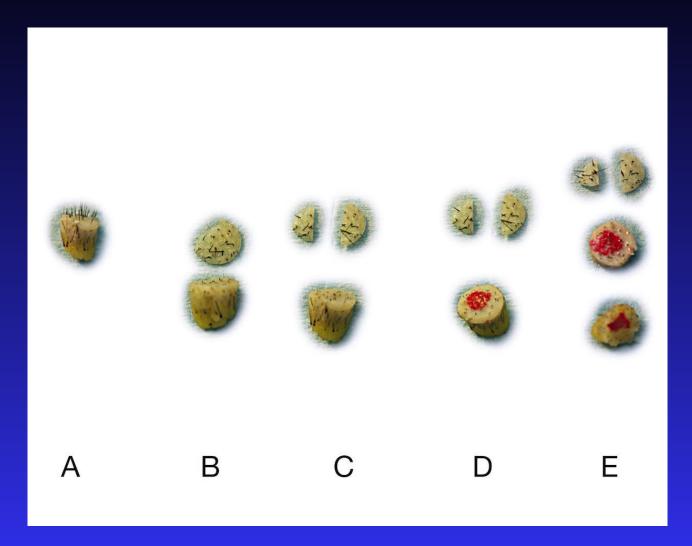
# Objective #1 Tissue Processing





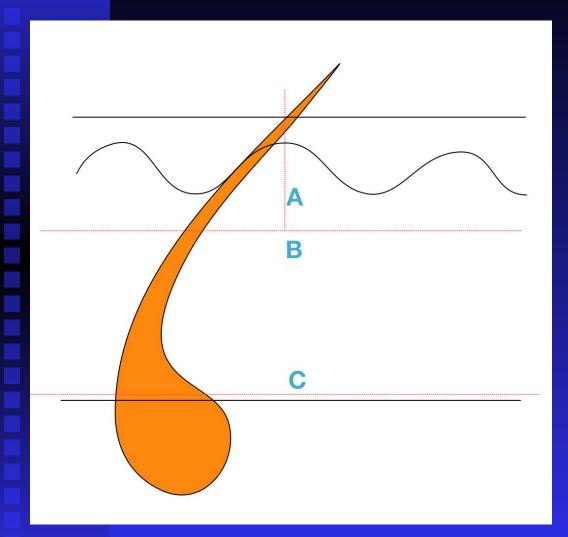


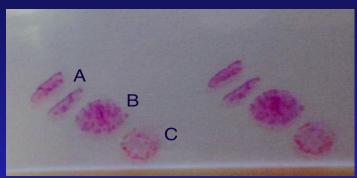
# Horizontal to Vertical = HoVert

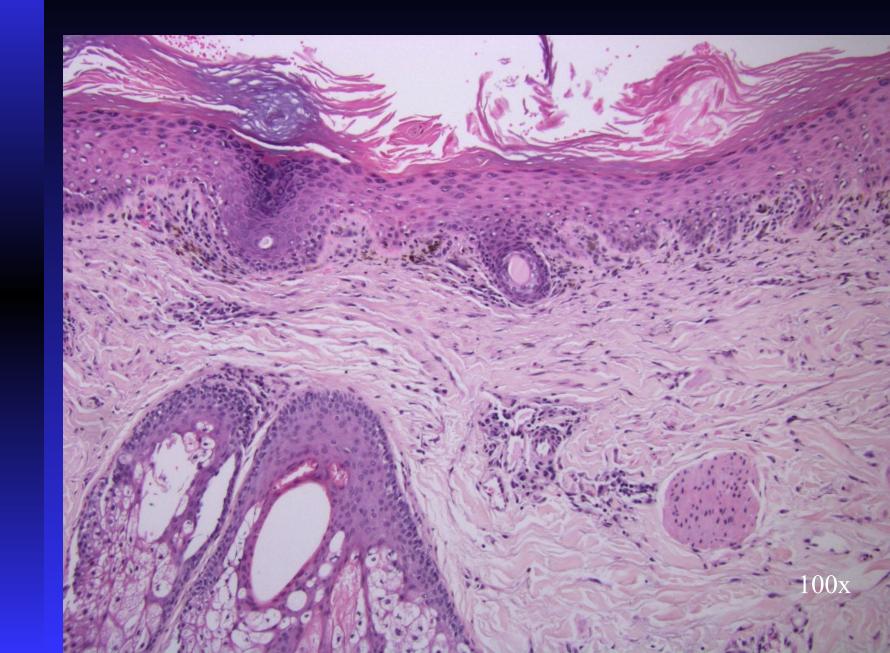


Nguyen JV, Hudacek K, Whitten JA, Rubin AI, Seykora JT. The HoVert technique: a novel method for the sectioning of alopecia biopsies. J Cutan Pathol 2011; 38: 401.

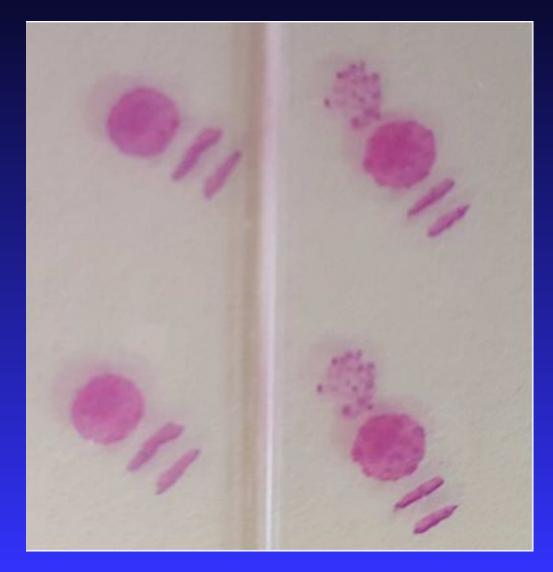
### Horizontal Vertical (HoVert) technique



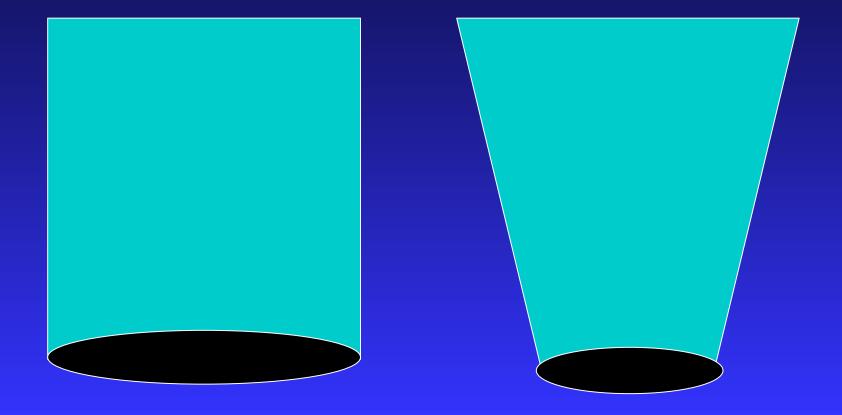




#### If biopsy is too thin, specimen bisected rather than trisected



# Not all 4mm punches the same



# Does not work well with transverse



# Objective #2 **H&E** Examination

### Follicular size

Vellus hairsShafts  $\leq$  inner root sheathShafts  $\leq$  0.03mm

Terminal hairs Shafts > inner root sheath Shafts  $\geq 0.06$ mm

# Telogen:vellus hair (T:V) ratio

> 3:1 Normal (<50 years old)</li>
 > 2:1 Normal (>50 years old):
 senescence
 < 2:1 Miniaturization</li>

Miniaturization Usually Only 2 diagnoses

 Female/male pattern hair loss (androgenetic)

Alopecia areata

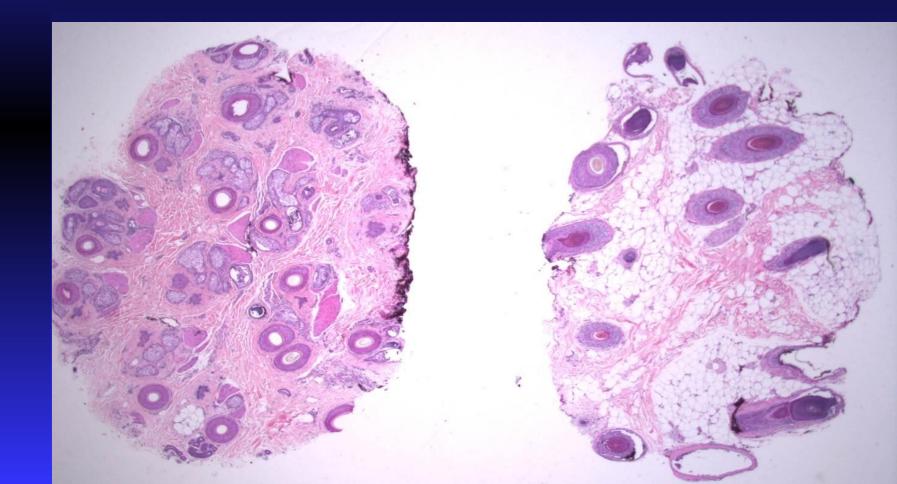
Female pattern hair loss vs Male pattern hair loss

- Female miniaturized to a certain point, then stops
- Male advances to complete loss of follicles

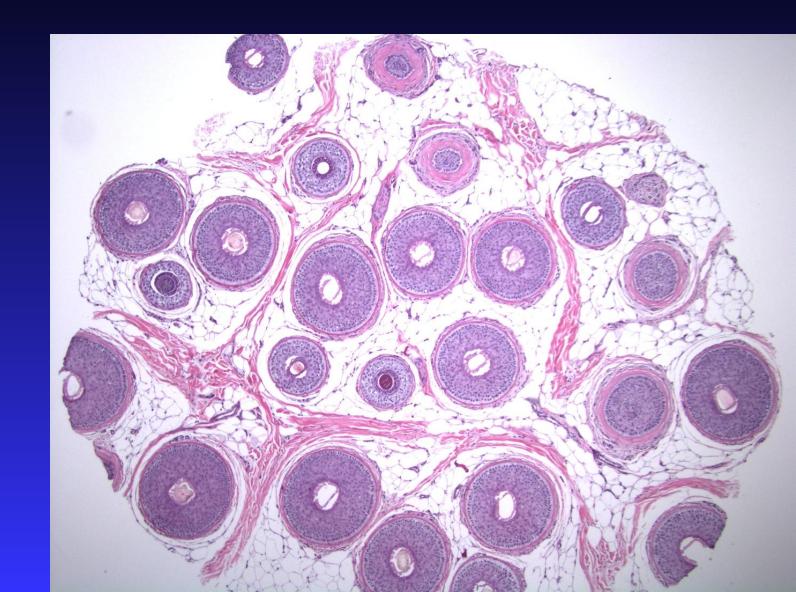
#### Intermediate Size—For diagnostic purposes group with vellus/miniaturizated

Indeterminate hairs 0.03mm < shafts < 0.06 mm

# Miniaturization

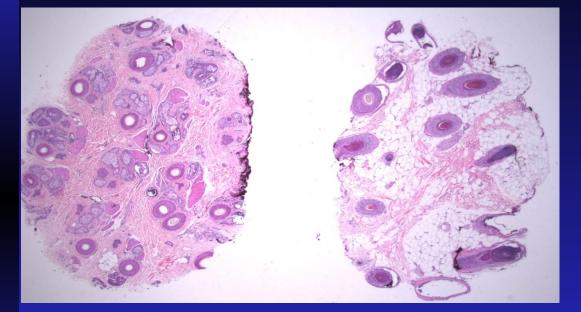


No miniaturization

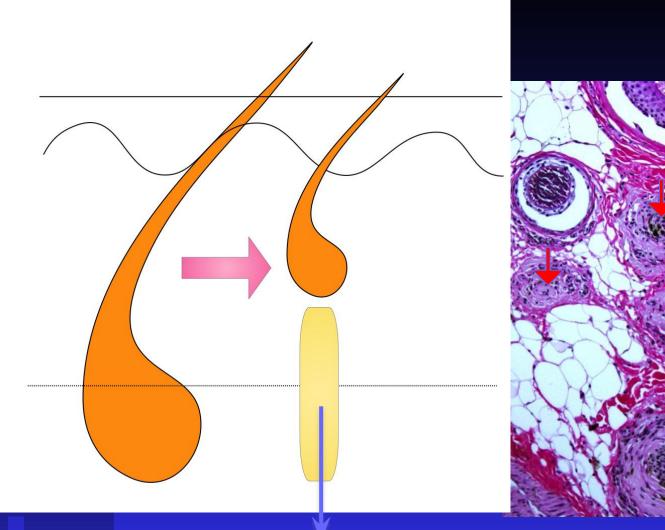


#### Total = Terminal + Vellus hairs

### **Terminal hairs**

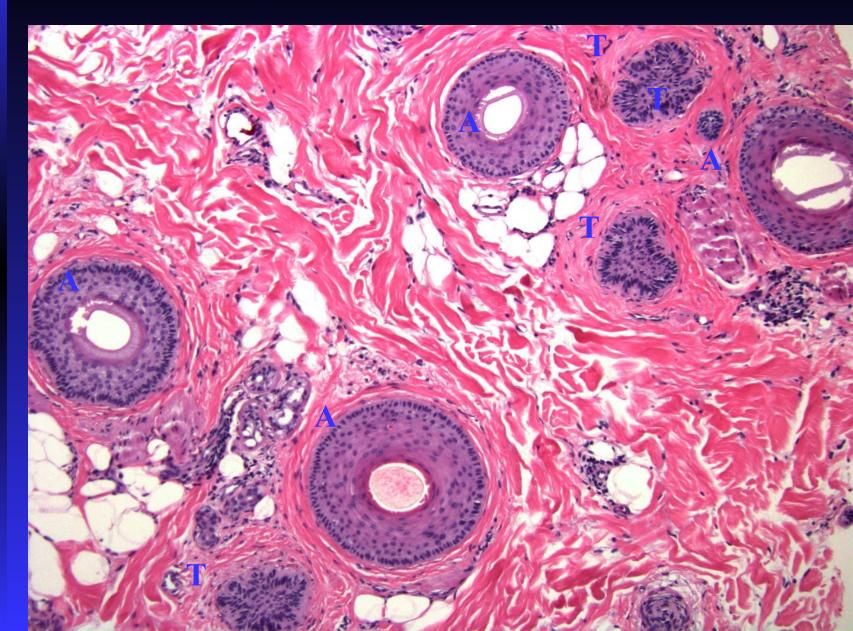


#### Total – Terminal = Vellus hairs

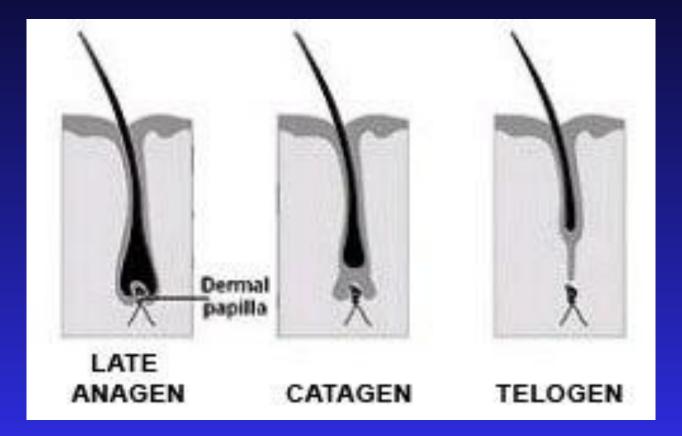


**Follicular fibrous tract (stela)** 

### Follicular Phase



### Normal Hair Cycle



http://www.hairsurgical.com/hair-loss/

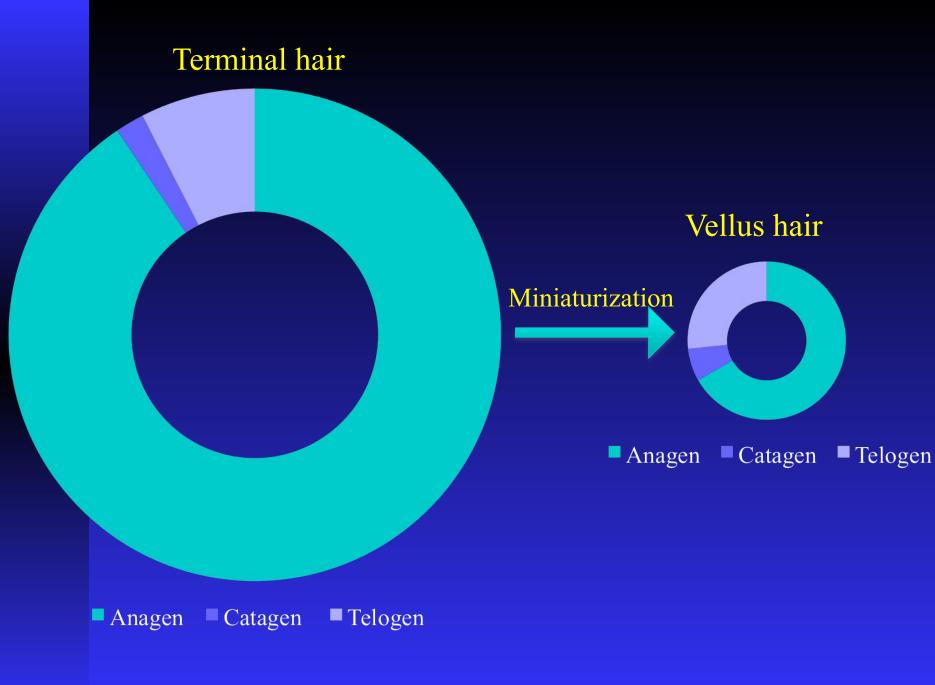
### Normal Hair Cycle—Scalp

- Anagen: >85% of all hairs; 5-7 years in length
- Catagen: <2%; 2 weeks</li>
  Telogen: <15%; 5 weeks</li>

#### Duration of Anagen Phase=Hair Length

Scalp 2-7 years

- Pubic/Axillary 3 months
- Eyebrows/eyelashes <1 month</p>



# Catagen-Telogen Importance in diagnosis

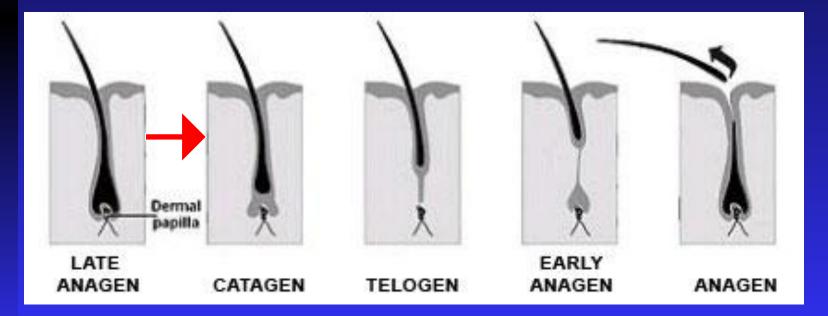
Acute Telogen Effluvium



Chronic Telogen Effluvium

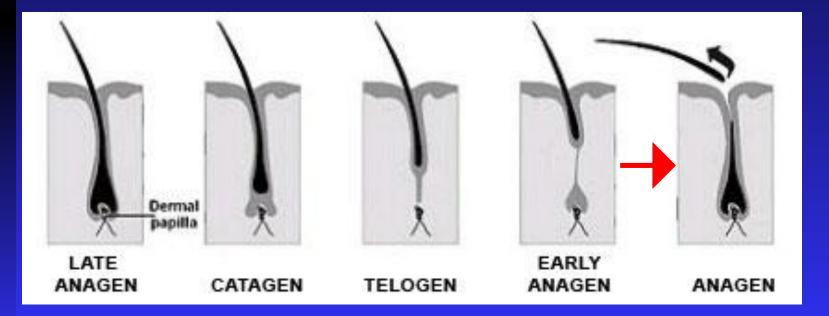
### Acute Telogen Effluvium.

 Profound, synchronized cycle into catagen/telogen

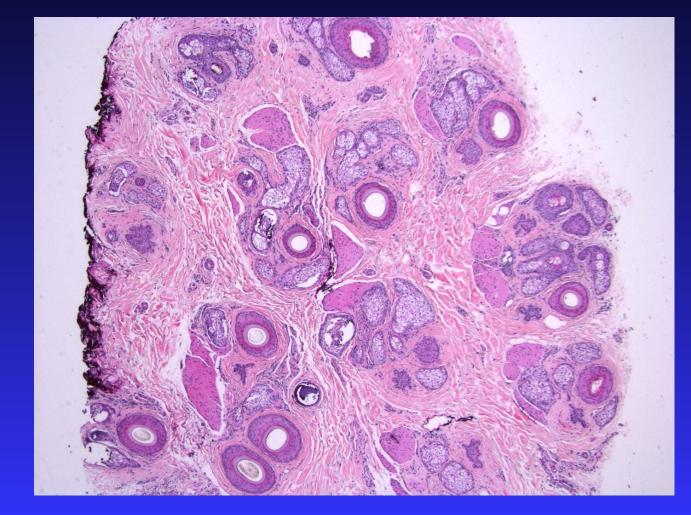


### Acute Telogen Effluvium.

 Shedding occurs several months after insult (fever, anesthesia, drug ingestion).



# Acute Telogen Effluvium. Right after insult

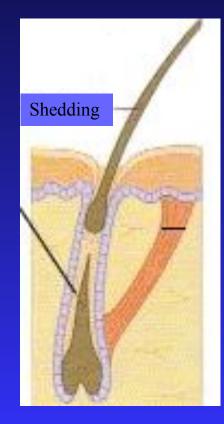


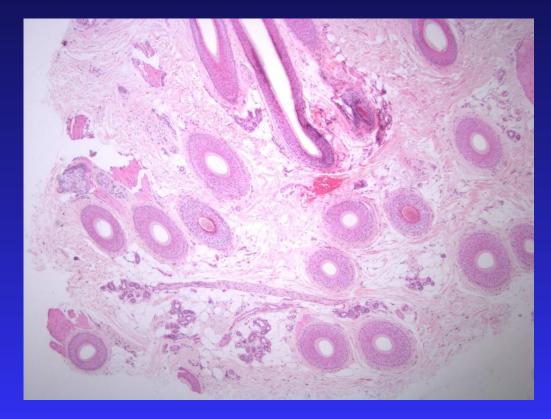
# Acute Telogen Effluvium. During shedding



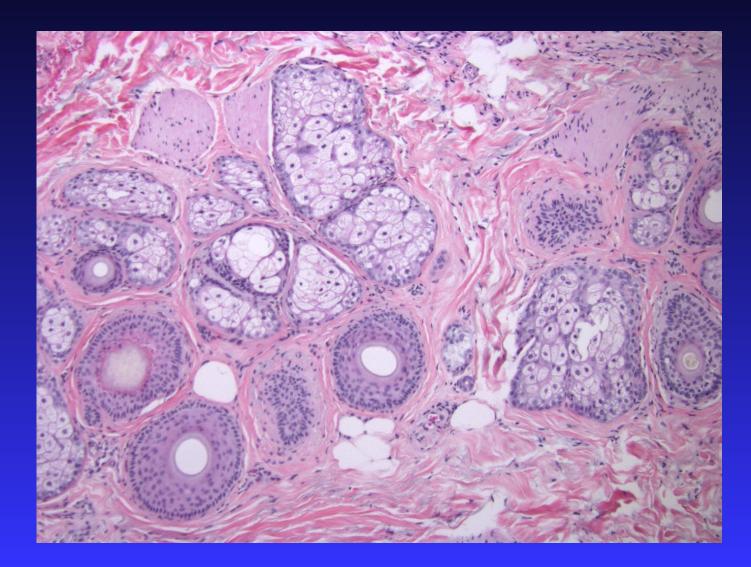
# Acute Telogen Effluvium

#### Histology: 100% Anagen phase follicles





#### Chronic Telogen Effluvium Histology: High catagen/telogen (>15-50%)



Acute Telogen Effluvium



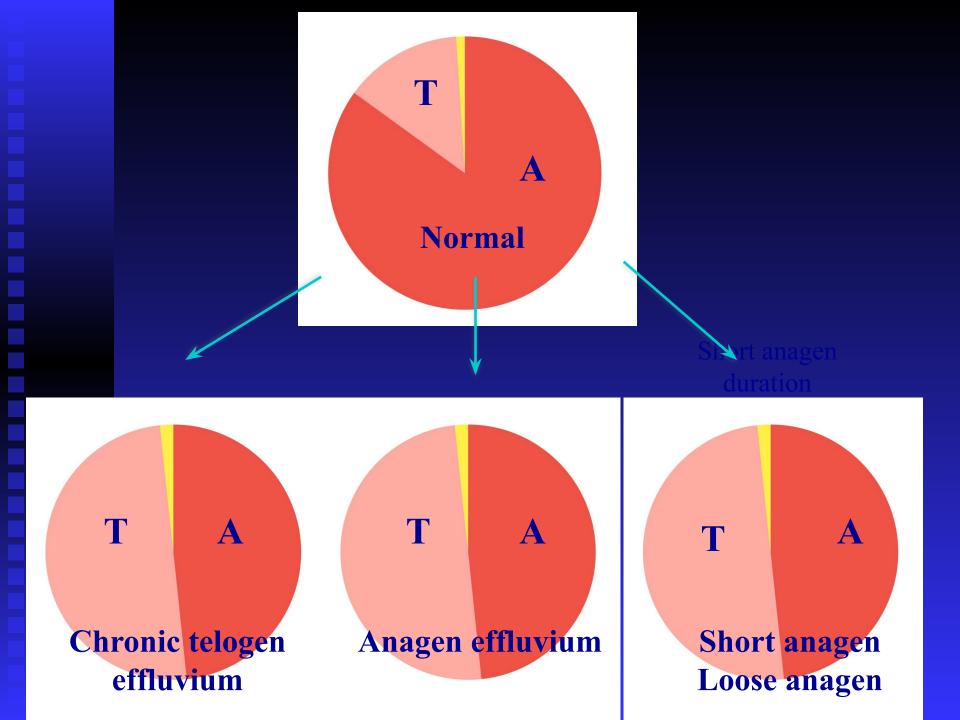
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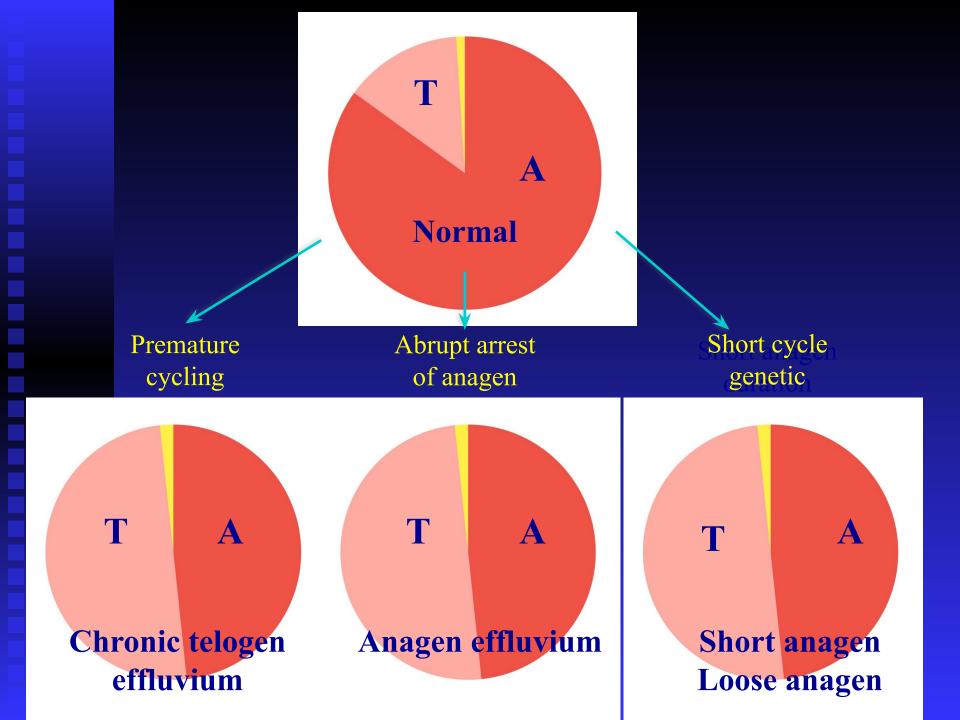


Chronic Telogen Effluvium



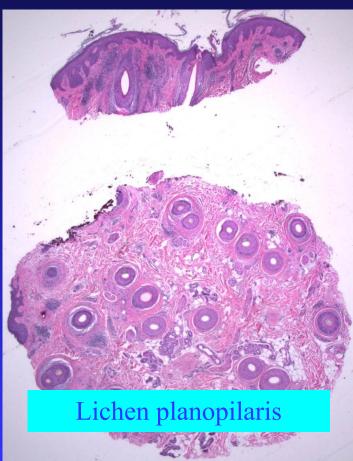
>20% Catagen/telogen





Diffuse alopecia with normal or absent catagen/telogen hairs: Acute telogen effluvium Diffuse lichen planopilaris





# Trichotillosis

TrichotillomaniaTraction alopecia

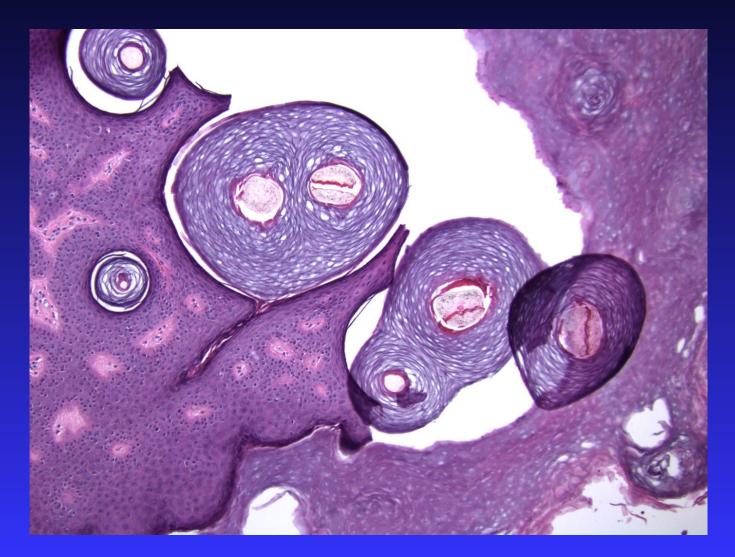
Trichotillosis Probably not good to group Trichotillomania Catagen/telogen shift Hair shaft breakage Melanin casts Lichen simplex chronicus Traction alopecia Low count (follicular loss) Catagen/telogen shift

## Trichotillomania: Hamburger Sign

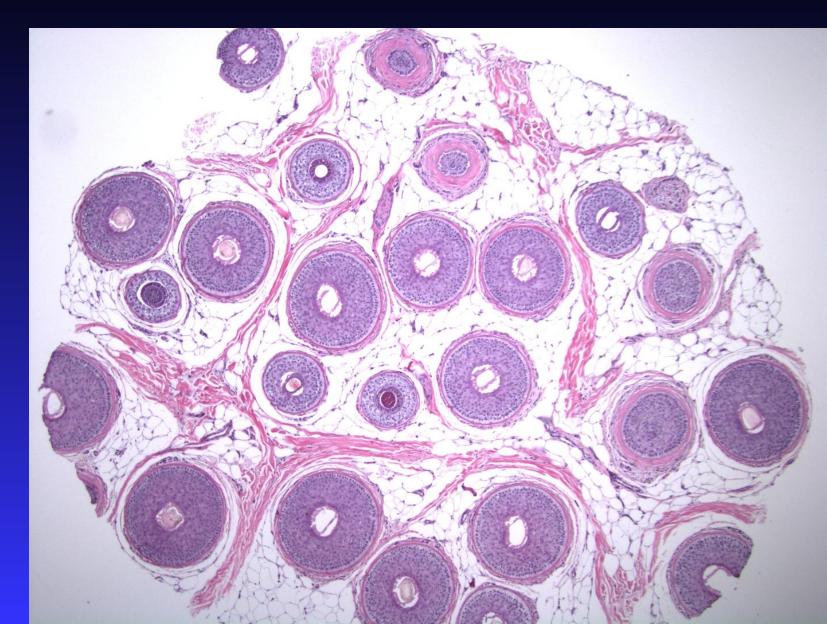
<u>Royer MC</u>Royer MC and <u>Sperling LC</u>Royer MC and Sperling LC. Splitting hairs: the 'hamburger sign' in <u>T</u>Royer MC and Sperling LC. Splitting hairs: the 'hamburger sign' in Trichotillomania. <u>J Cutan Pathol.</u> 33Suppl 2:63-4, 2006.



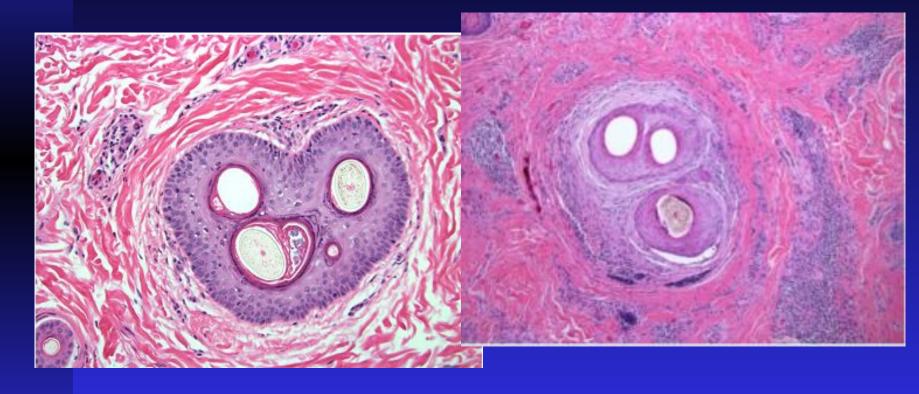
## Lichen simplex chronicus



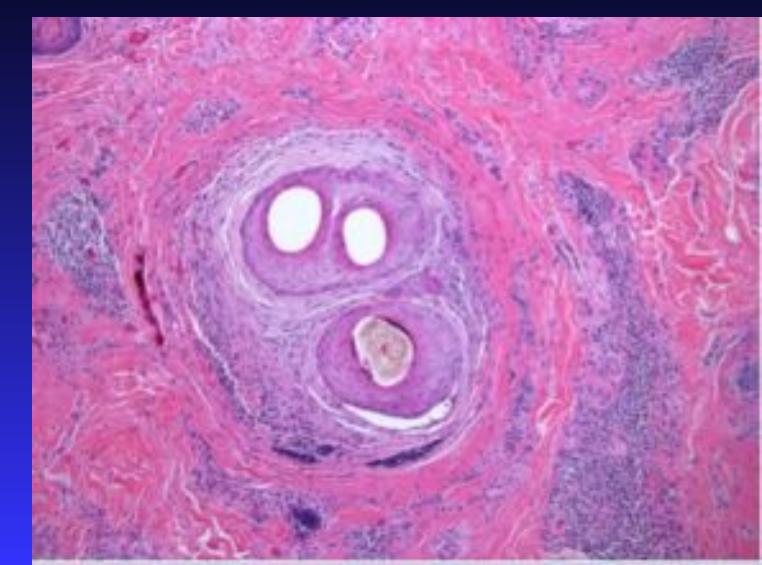
## Trichotillosis-Catagen/Telogen shift



## Normal Ostium—Not scar



# May be a resolving acneiform lesion (Zit)



## Lichen Planopilaris (LPP)

## Miniepidemic?

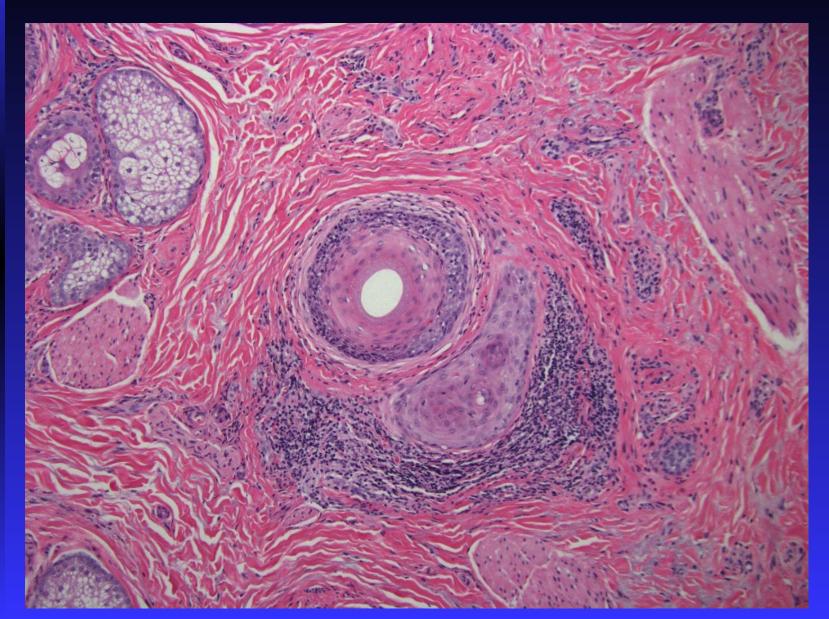
- Hair loss clinicians observing increased incidence.
- Traditionally USWest Coast > East Coast
- Observed during exams when patient being seen for other problem

Lichen Planopilaris Increasing Incidence

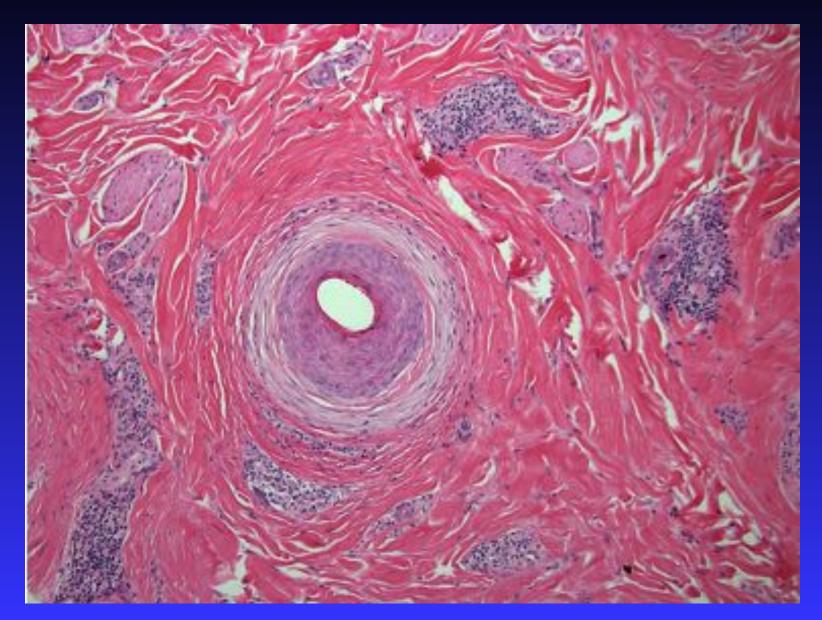
- Nanoparticle
  - Lichen planus—Metals, especially dental implicated
    - Gold, mercury—dental\*
    - Nail LP associated with +metal patch test\*\*

\*Sasaki G et al. J Dermatol 23:890, 1996. \*\*Nishizawa A et al. J Eur Acad Dermatol Venerol 27:e231, 2013.

# Lichen Planopilaris



# Lichen Planopilaris



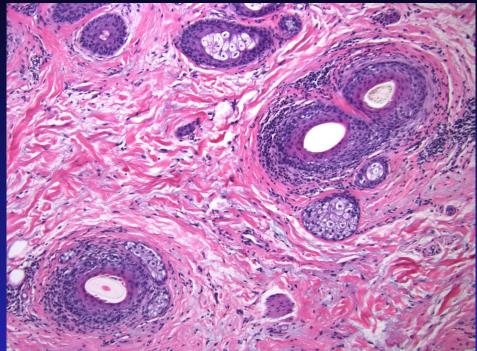
## LPP—Frontal Fibrosing Variant

 Appears to target smaller follicles (eyebrows, vellus/body)
 Increasing incidence

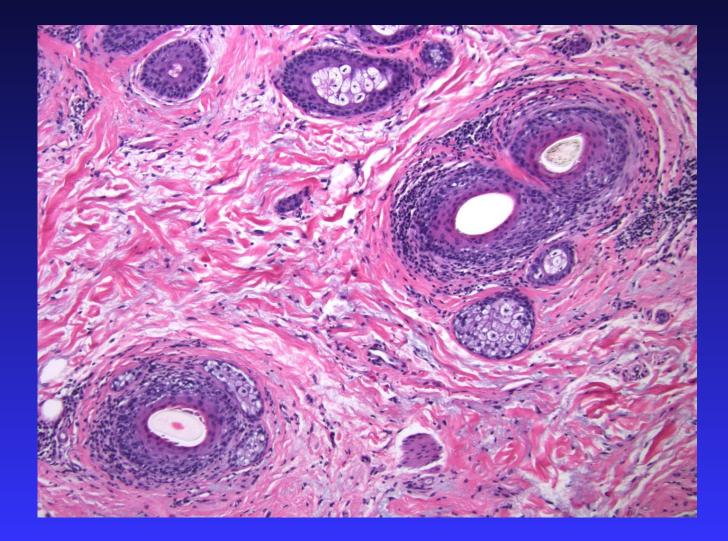


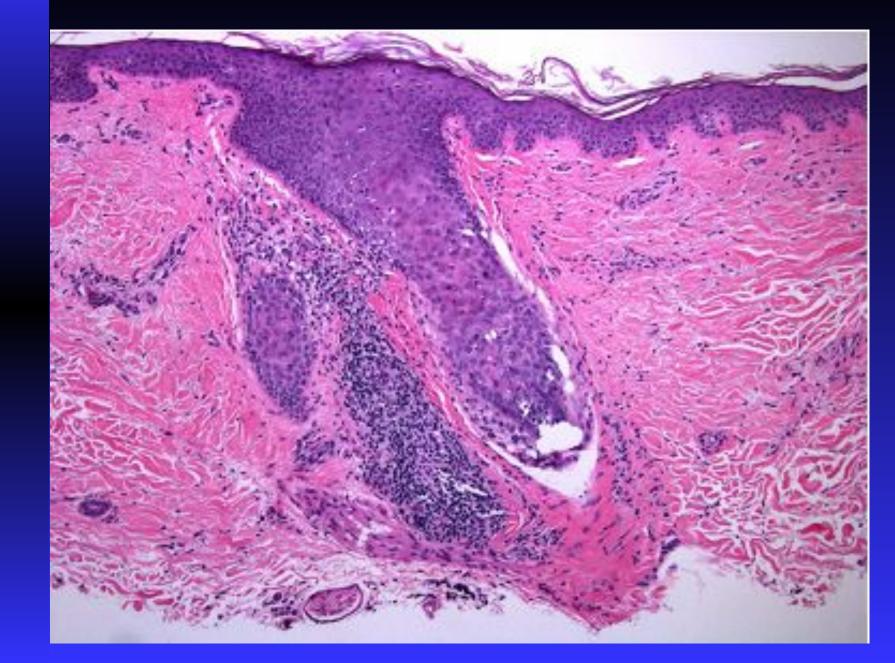


## Frontal fibrosing alopecia

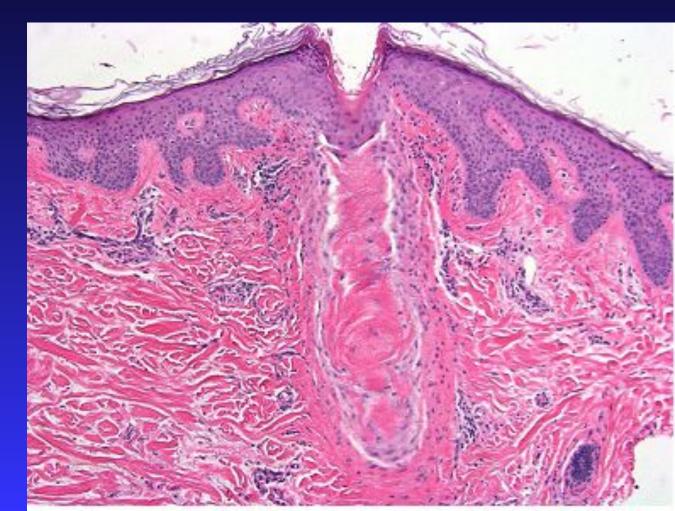


## LPP—Frontal Fibrosing Variant Histology: Minimal or no scarring





LPP Frontal Fibrosing--Eyebrow Often complete follicular loss with lymphocytes and no scarring



## Concept

All alopecias, whether classified as cicatricial or non-cicatricial, may, in the end, be scarring

	Cicatricial alopecia	Non-cicatricial alopecia
Follicular and sebaceous epithelium	Destroyed	Preserved
Final outcome	Replacement by follicular scarring	Regrowth
Hair loss	Irreversible	Reversible

# **Not Necessarily True**

	Cicatricial alopecia	Non-cicatricial alopecia
Follicular and sebaceous epithelium	Destroyed	Preserved
Final outcome	Replacement by follicular scarring	Regrowth
Hair loss	Irreversible	Reversible

## **Biphasic alopecia**

Long standing alopecia Follicular stem cell exhaustion

Follicular drop out Empty fibrous tract



Sperling LC, Lupton GP. Histopathology of non-scarring alopecia. J Cutan Pathol 1995; 22: 97-114.

# Objective #3

## Immunohistochemistry for diagnostic impasses



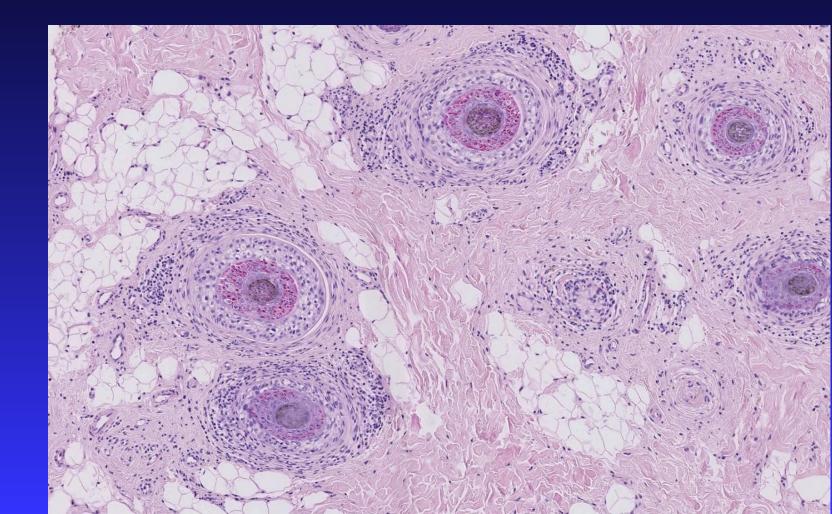


# 





## Acute alopecia areata « Hive of Bees »



# Alopecia areata incognita



Courtesy BM Piraccini, Bologna

	Pattern hair loss	Alopecia areata
Terminal/Vellus hair ratio	Decrease	Decreased
% catagen and telogen hair	Increased	Increased

- 1. Elston DM, Ferringer T, Dalton S, Fillman E, Tyler W. A comparison of vertical versus transverse sections in the evaluation of alopecia biopsy specimens. J Am Acad Dermatol 2005; 53: 267.
- 2. Wohltmann WE, Sperling LC. Histopathologic diagnosis of multifactorial alopecia. J Cutan Pathol 2016; 43: 483.

How to make the distinction between alopecia areata and pattern hair loss in absence of peribulbar lymphocytic infiltrate?

## ARTICLE IN PRESS

### DERMATOPATHOLOGY

## Distinguishing diffuse alopecia areata (AA) from pattern hair loss (PHL) using CD3<sup>+</sup> T cells

Athanassios Kolivras, MD,<sup>a</sup> and Curtis Thompson, MD<sup>b</sup> Brussels, Belgium, and Portland, Oregon

**Background:** Distinguishing between diffuse subacute alopecia areata (AA), in which the peribulbar infiltrate is absent, and pattern hair loss is challenging, particularly in cases that lack marked follicular miniaturization and a marked catagen/telogen shift.

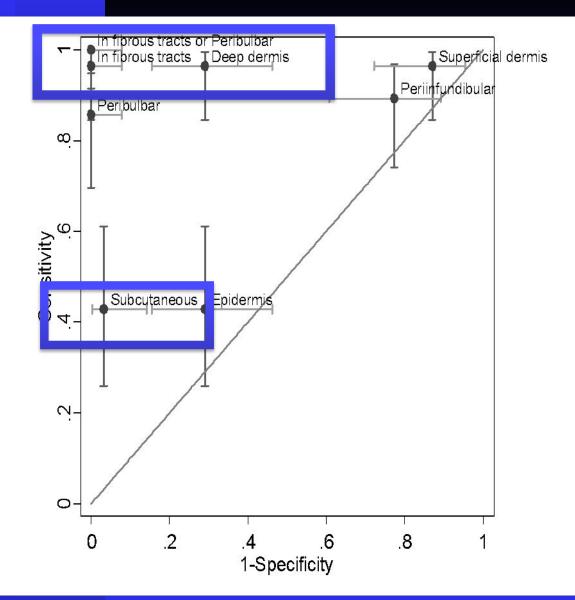
**Objective:** We sought to distinguish diffuse AA from pattern hair loss using CD3<sup>+</sup> T lymphocytes.

Metbods: A total of 28 cases of subacute AA and 31 cases of pattern hair loss were selected and a 4-mm punch biopsy was performed. All the specimens were processed using the "HoVert" (horizontal and vertical) technique. In all cases, hematoxylin-eosin and immunohistochemical stains for CD3, CD4, CD8, and CD20 were performed.

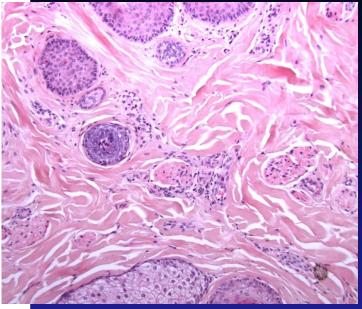
**Results:** The presence of CD3<sup>+</sup> lymphocytes within empty follicular fibrous tracts (stela), even without a concomitant peribulbar infiltrate, is a reliable histopathological clue in supporting a diagnosis of AA (sensitivity 0.964, specificity 1,  $P \leq .001$ ).

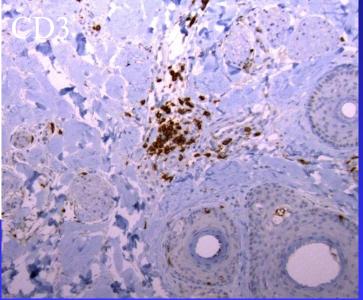
Limitations: Limited tissue for analysis remained in the clinical sample tissue blocks.

**Conclusion:** The presence of CD3<sup>+</sup> T-cells within empty follicular fibrous tracts (stela) supports a diagnosis of AA. (J Am Acad Dermatol http://dx.doi.org/10.1016/j.jaad.2015.12.011.)

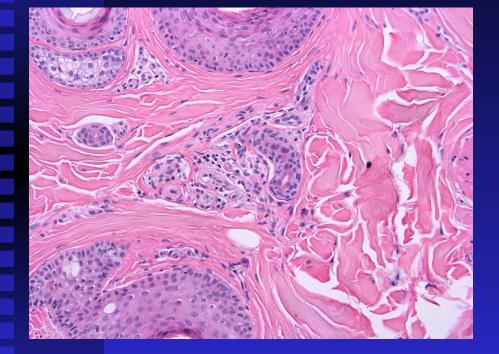


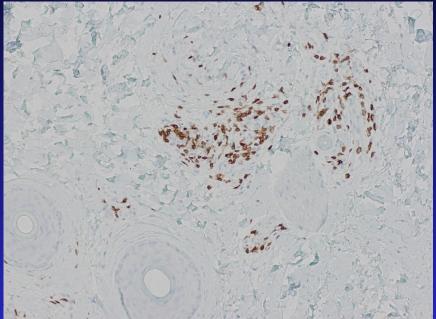
## Alopecia areata





# AA = CD3 in empty tracts





## CONCLUSION

Presence of CD3+ T-lymphocytes within the empty fibrous follicular tracts favors a diagnosis of alopecia areata



## Courtesy BM Piraccini, Bologna

## Lichen planopilaris or Discoid lupus erythematosus ?

#### LPP





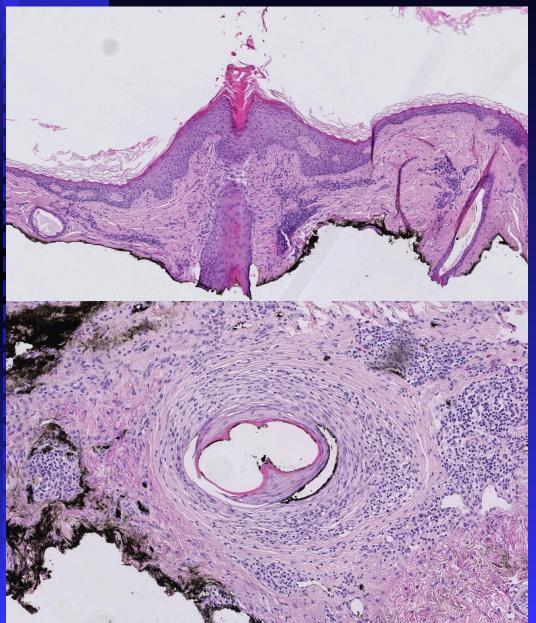






Courtesy BM Piraccini, Bologna

# LPP

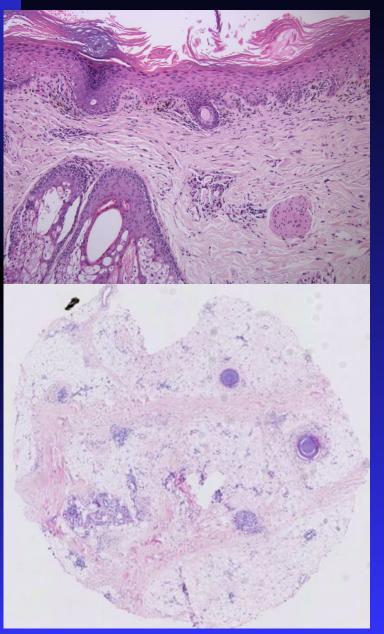


1. Perifollicular scarring and lymphocytic infiltrate at the level of the infundibulum or superficial isthmus

# 2. No inter-follicular epidermal interface dermatitis

#### 3. No deep infiltrate

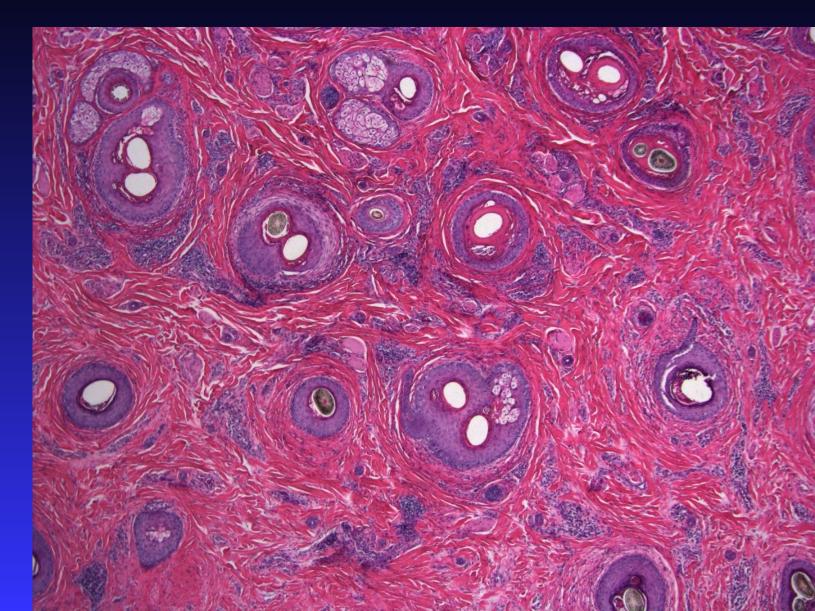
# LE



# Interface change in the interfollicular epidermis

#### Deep dermal or subcutaneous lymphocytes

# LPP or LE?



#### Lupus erythematosus IgG

### Plasmacytoid dendritic cells: CD123

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Journal of Cutaneous Pathology

J Cutan Pathol 2010: 37: 1132-1139 doi: 10.11111/j.1600-0560.2010.01587.x John Wiley & Sons. Printed in Singapore



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Plasmacytoid dendritic cells: an overview of their presence and distribution in different inflammatory skin diseases, with special emphasis on Jessner's lymphocytic infiltrate of the skin and cutaneous lupus erythematosus Dario Tomasini<sup>1</sup>, Thomas Mentzel<sup>2</sup>, Markus Hantschke<sup>2</sup>, Amilcare Cerri<sup>3</sup>, Bruno Paredes<sup>2</sup>, Arno Rütten<sup>2</sup>, Leo Schärer<sup>2</sup> and Heinz Kutzner<sup>2</sup>

<sup>1</sup>Department of Dermatology, Hospital of Busto Arsizio, Busto Arsizio, Italy, <sup>2</sup>Dermatopathologie Friedrichshafen, Friedrichshafen, Germany, and <sup>3</sup>Department of Medicine, Surgery and Dentistry, University of Milan, Milan, Italy

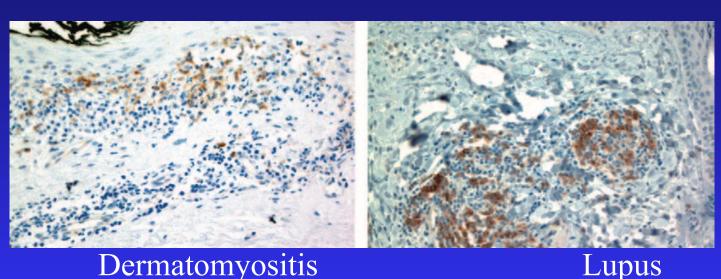


J Cutan Pathol 2008: 35: 452–456 doi: 10.1111/j.1600-0560.2007.00848.x Blackwell Munksgaard. Printed in Singapore

Plasmacytoid dendritic cells are present in cutaneous dermatomyositis lesions in a pattern distinct from lupus erythematosus

#### Jennifer M. McNiff<sup>1,2</sup> and Daniel H. Kaplan<sup>3</sup>

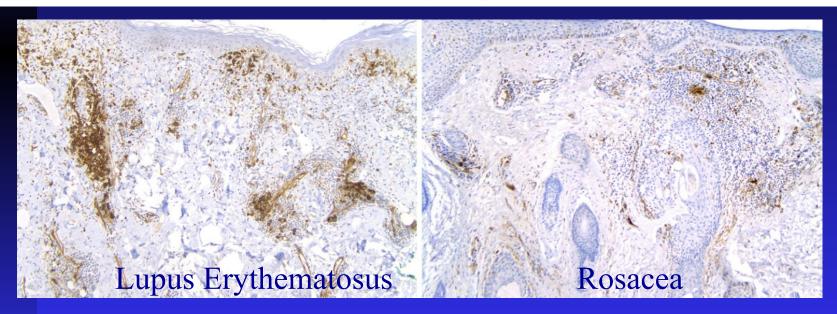
<sup>1</sup>Department of Dermatology and <sup>2</sup>Department of Pathology, Yale University School of Medicine, New Haven, CT, USA <sup>3</sup>Department of Dermatology, University of Minnesota, Minneapolis, MN, USA



#### Dermatomyositis Erythematosus

#### Comparative analysis of rosacea and cutaneous lupus erythematosus: Histopathologic features, T-cell subsets, and plasmacytoid dendritic cells

Theodore T. Brown, MD,<sup>a</sup> Eun-Young K. Choi, MD,<sup>a</sup> Dafydd G. Thomas, MD, PhD,<sup>a</sup> Alexandra C. Hristov, MD,<sup>a,b</sup> and May P. Chan, MD<sup>a,b</sup> Ann Arbor, Michigan



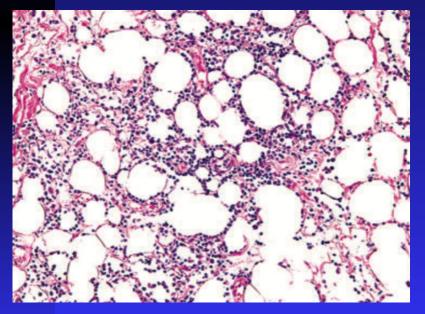
*Conclusion:* The infiltrates in rosacea and LE differ immunophenotypically, and may aid in their distinction in addition to conventional histologic examination. (J Am Acad Dermatol 2014;71:100-7.)

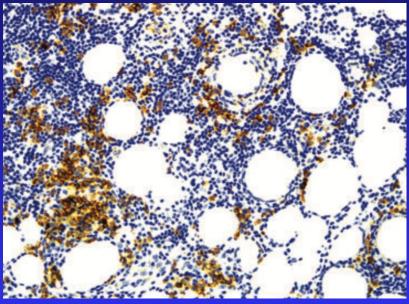
Histopathology 2013, 62, 1057-1066. DOI: 10.1111/his.12105



#### The presence of clusters of plasmacytoid dendritic cells is a helpful feature for differentiating lupus panniculitis from subcutaneous panniculitis-like T-cell lymphoma

Jau-Yu Liau,<sup>1</sup> Shih-Sung Chuang,<sup>2</sup> Chia-Yu Chu,<sup>3</sup> Wen-Hui Ku,<sup>4</sup> Jia-Huei Tsai<sup>1</sup> & Teng-Fu Shih<sup>5</sup>



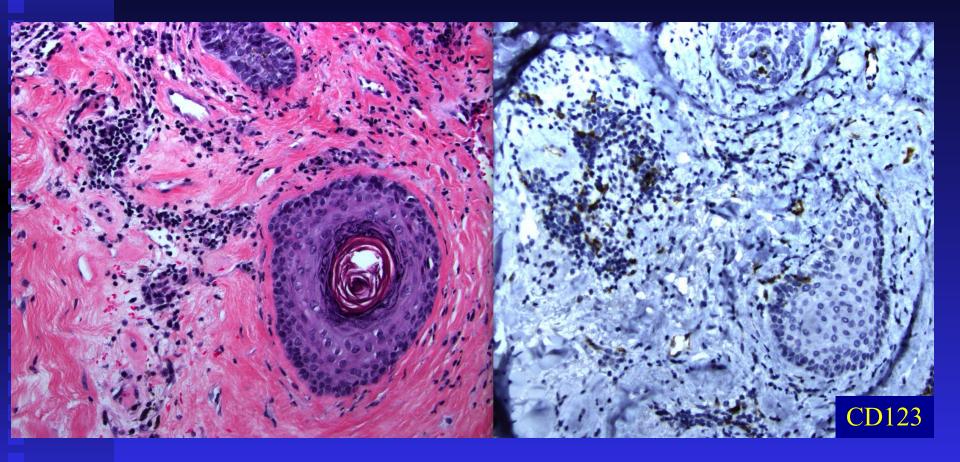


Subcutaneous panniculitis-like T-cell lymphoma

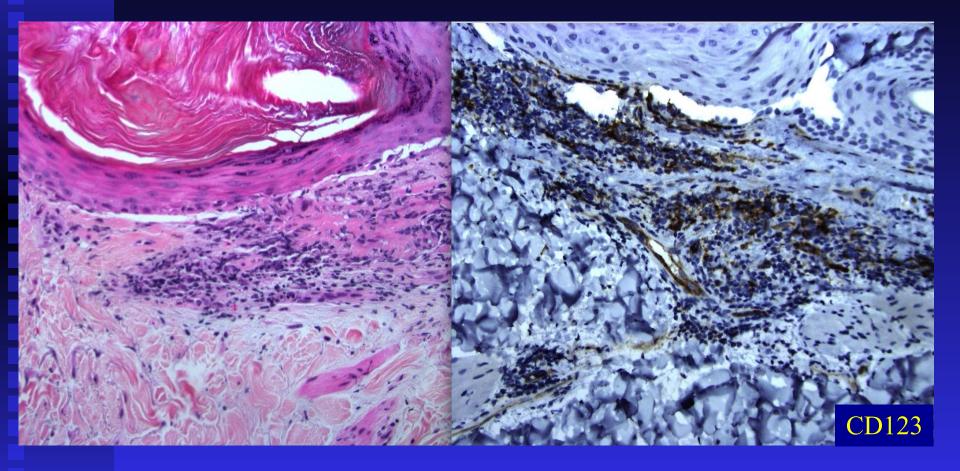
Kolivras A, Thompson C. Clusters of CD123+ plasmacytoid dendritic cells help distinguish lupus alopecia from lichen planopilaris. J Am Acad Dermatol 2016; 74: 1267–69.



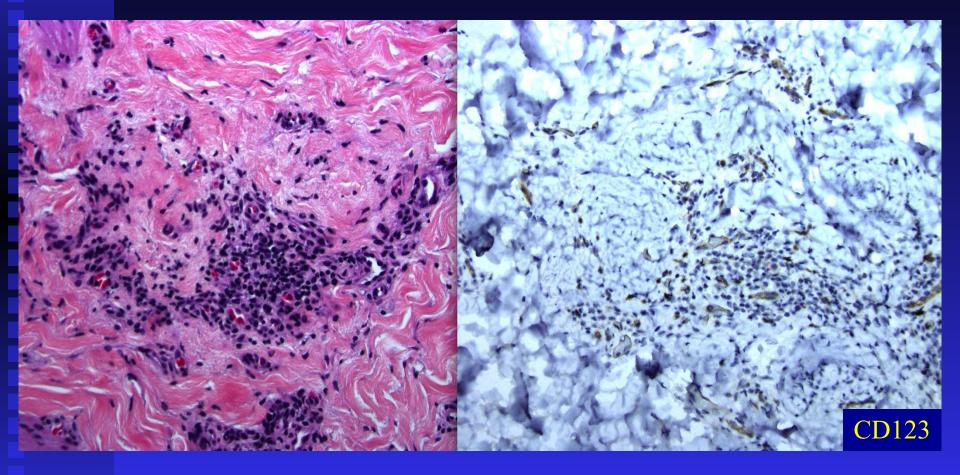
# Diffuse dermal expression



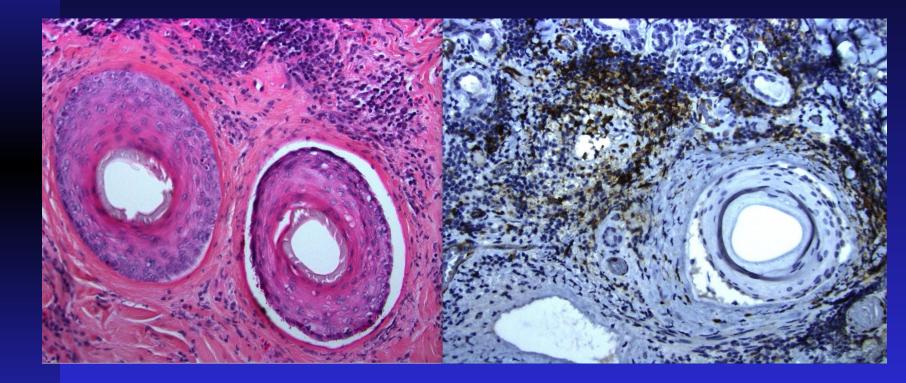
# Underneath perifollicular epidermis

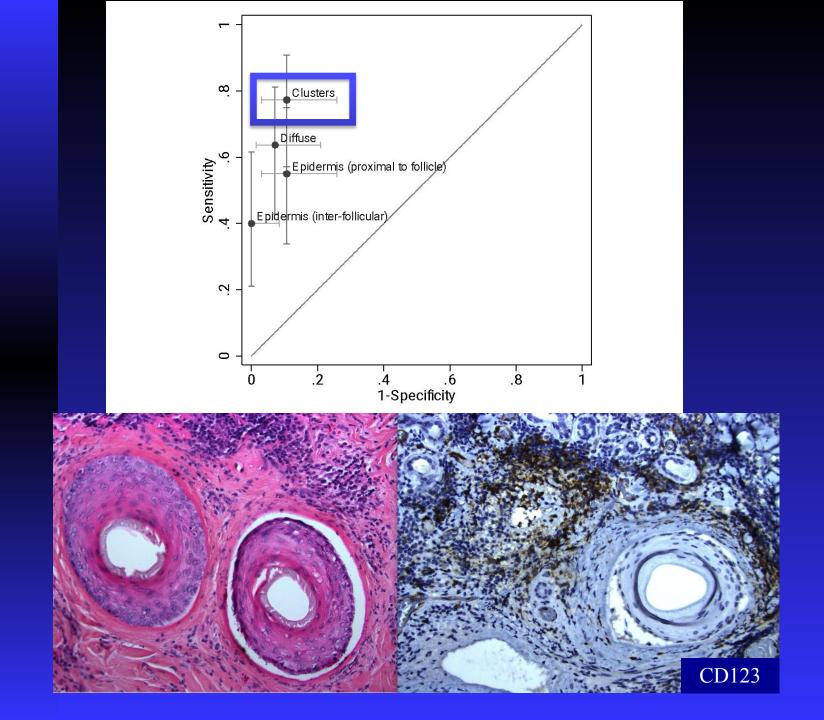


## Underneath interfollicular epidermis

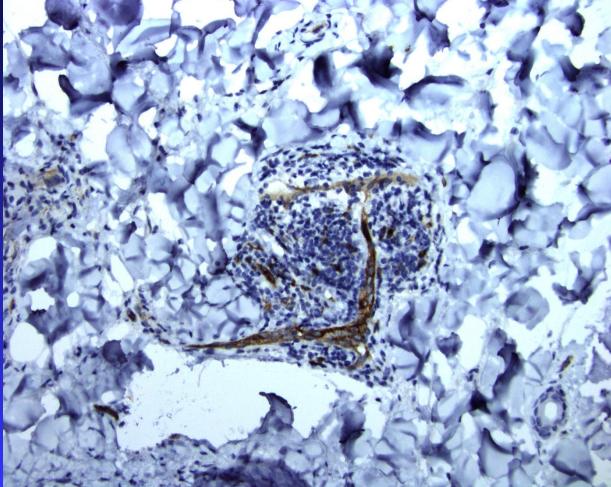


# Clusters of CD123 = Lupus





Warning: CD123 positive in endothelium



# CONCLUSION

**Clusters of CD123+ plasmacytoid dendritic cells favor a diagnosis of lupus erythematosus**  CD123 immunohistochemistry for plasmacytoid dendritic cells is useful in the diagnosis of scarring alopecia

J Cutan Pathol 2016; 43: 643-8

#### Katherine Fening<sup>1</sup>, Viswas Parekh<sup>2</sup> and Kristopher McKay<sup>1</sup>

<sup>1</sup>Division of Dermatopathology, University of Alabama at Birmingham, Birmingham, AL, USA and <sup>2</sup>Department of Pathology, University of Alabama at Birmingham, Birmingham, AL, USA

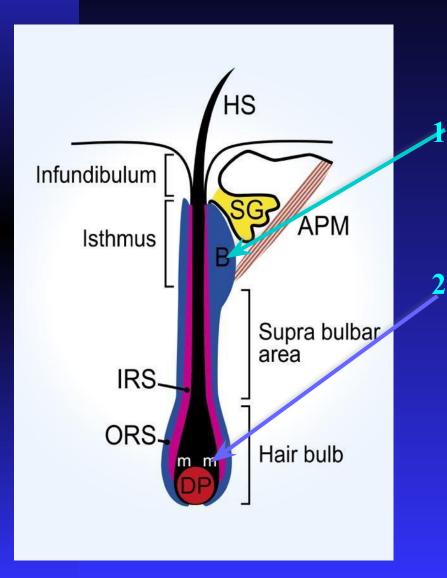
# Lichen planopilaris Lupus erythematosus

# Lichen Planopilaris (LPP)

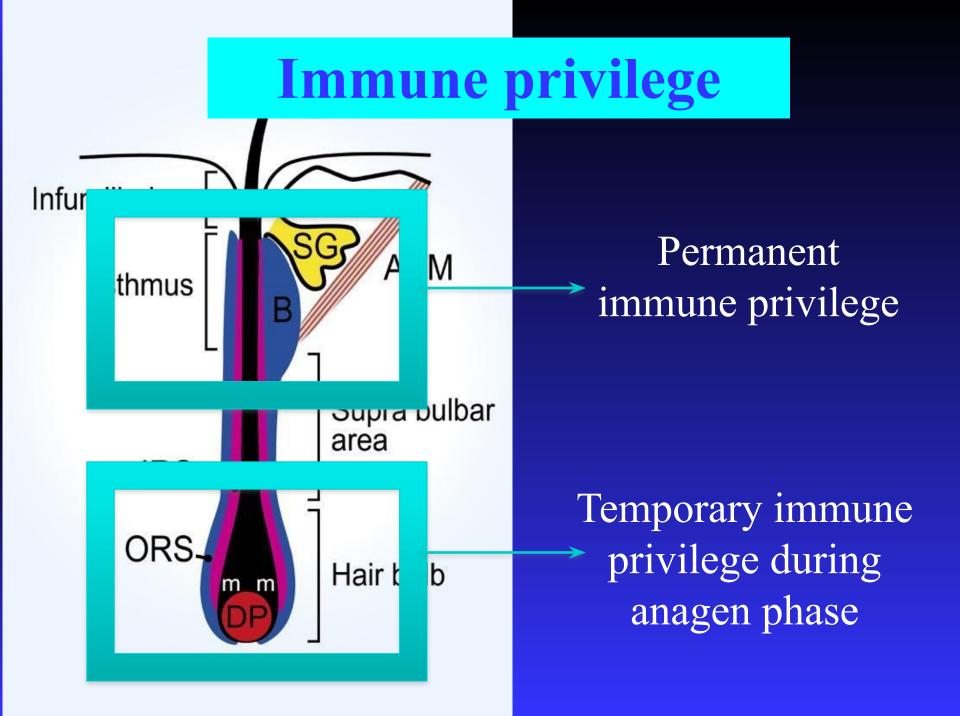
"Both the patient and the doctor ask the vexing question: "When will it end?"... LPP sometimes has no apparent ending."

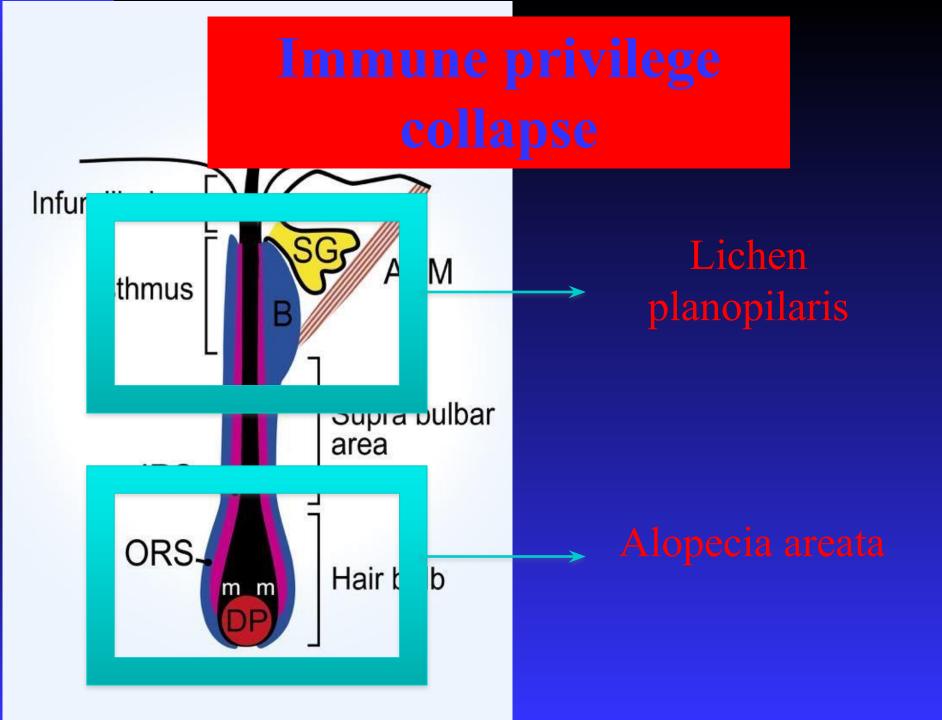
Philippe Assouly and Pascal Reygagne. Lichen Planopilaris: Update on Diagnosis and Treatment. Semin Cutan Med Surg 28:3-10, 2009.

## Two Stem Cells



Bulge stem cells:
quiescent and maintain
long-term stem cell pool
Hair germ cells:
activated during
anaphase engaging in a
new growth





**Cutaneous** Pathology

Possible role of the bulge region in the pathogenesis of inflammatory scarring alopecia: lichen planopilaris as the prototype

#### Narciss Mobini, Sam Tam and Hideko Kamino

Department of Dermatology, Section of Dermatopathology, New York University Medical Center, New York, NY, USA

**Conclusion:** Our study supports the finding that in LPP, the inflammatory infiltrate mainly involves the bulge region, where the stem cells reside. Once this area is damaged, the hair loses its potential of regrowth with resulting scarring alopecia. This is in contrast with inflammatory non-scarring alopecias such as alopecia areata, where the bulb region is targeted, sparing the stem cells.

Journal of Pathology J Pathol 2013; 231: 236–247 Published online in Wiley Online Library (wileyonlinelibrary.com) DOI: 10.1002/path.4233

#### Lichen planopilaris is characterized by immune privilege collapse of the hair follicle's epithelial stem cell niche

Matthew J Harries,<sup>1</sup> Katja Meyer,<sup>2</sup> Iskander Chaudhry,<sup>3</sup> Jennifer E Kloepper,<sup>2</sup> Enrique Poblet,<sup>4</sup> Christopher EM Griffiths<sup>1</sup> and Ralf Paus<sup>1,2</sup>\*

<sup>1</sup> Dermatology Centre, Salford Royal NHS Foundation Trust, University of Manchester, Manchester Academic Health Science Centre, UK

- <sup>2</sup> Department of Dermatology, University of Lübeck, Germany
- <sup>3</sup> Department of Pathology, Central Manchester NHS Foundation Trust, UK

<sup>4</sup> Department of Pathology, University General Hospital of Murcia, Spain

\*Correspondence to: R Paus, Department of Dermatology, University of Lübeck, Lübeck, Germany. E-mail: ralf.paus@uksh.de or ralf.paus@manchester.ac.uk

"These novel findings raise the possibility that LPP represents an autoimmune disease in whose pathogenesis **IFNy-induced bulge immune privilege collapse** plays an important role."

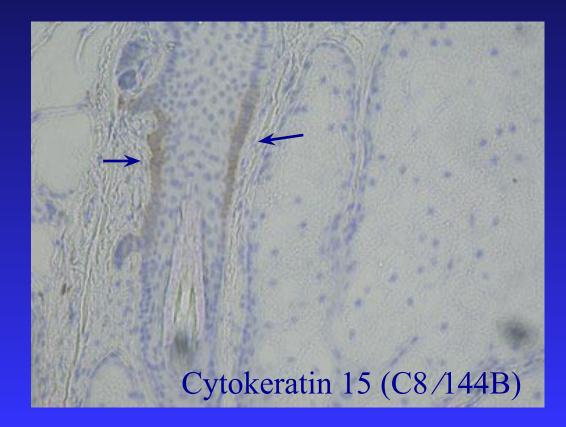
#### **REVIEW ARTICLE**

#### Stem cells and alopecia: a review of pathogenesis

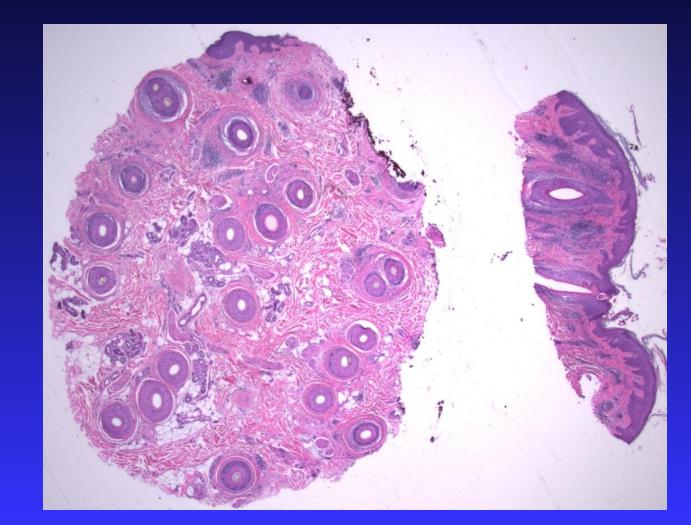
K. Al-Refu

Faculty of Medicine, Internal Medicine Department, Mu'tah University, PO Box 5, Karak, Mu'tah 61710, Jordan

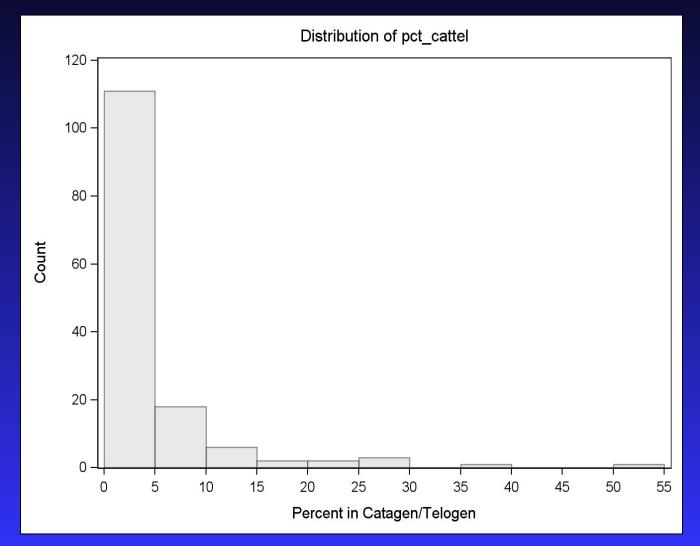
Br J Dermatol 2012; 167: 479-484.



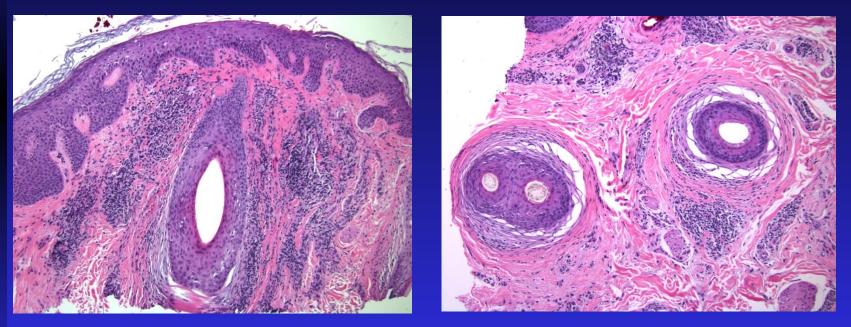
# Near absence of catagen/telogen in lichen planopilaris



## Near absence of catagen/telogen in LPP

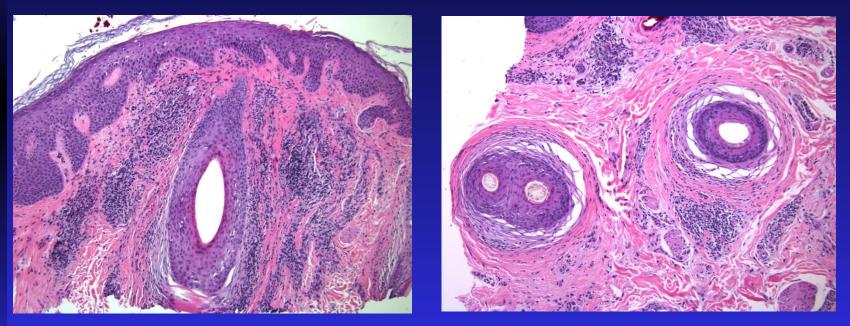


# Lichen Planopilaris (LPP)



#### Inflammation at level of the follicular bulge.

# Lichen Planopilaris (LPP)

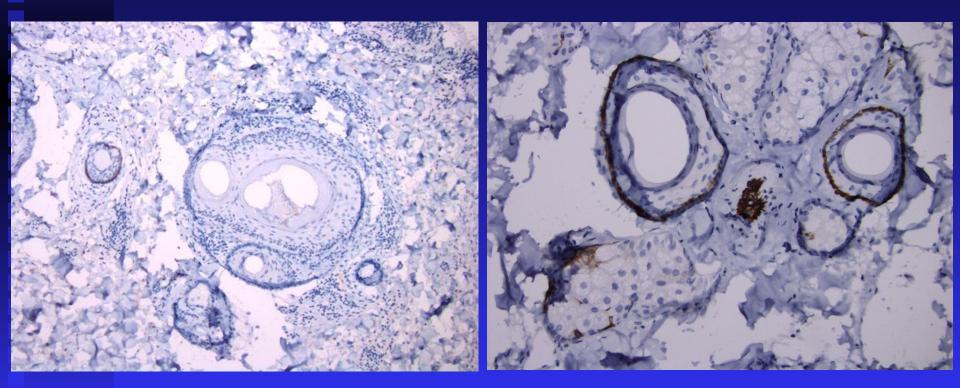


Loss of catagen/telogen phase follicles

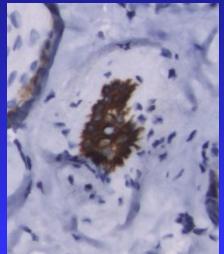
#### Absence of catagen/telogen phase and loss of cytokeratin 15 expression in hair follicles in lichen planopilaris

Arlette Habashi-Daniel, MD,<sup>a</sup> Janet L. Roberts, MD,<sup>b</sup> Nisha Desai, MD,<sup>d</sup> and Curtis T. Thompson, MD<sup>a,b,c</sup> *Portland, Oregon* 

J Am Acad Dermatol 2014;71:969-72.



# Lichen Planopilaris Progression Loss of CK15+ stem cells Disappearance of follicle when cycle into catagen



Clinical progression despite immunosuppressive treatment

## Lichen Planopilaris (LPP)

Absence of catagen/telogen

Loss of CK15+ stem cells

EARLY INTERVENTION IS LIKELY THE MOST IMPORTANT FACTOR IN PROGNOSIS.

## CK15+ Stem Cell Loss in Other Cicatricial Diseases

- Alopetic lupus erythematosus—(7 cases)
- Central centrifugal cicatricial alopecia (CCCA)—(7)
- Frontal fibrosing alopecia (LPP)—(9)
  Lichen planopilaris (19)

Kolivras A, Thompson N, Thompson C. Loss of cytokeratin-15 (CK15) expression is not specific for lichen planopilaris (LPP). J Am Acad Dermatol 2016; 75: 428-9.

## CONCLUSION

Loss of CK15+ follicular bulge cells has no diagnostic value in distinguishing cicatricial alopecias



FRIMART CICATRICIAL ALOFECIA	PRIMARY	<b>CICATRICIAL</b>	<b>ALOPECIA</b>
------------------------------	---------	--------------------	-----------------

#### Lymphocytic Chronic cutaneous lupus erythematosus

Lichen planopilaris (LPP)

- Classic LPP

- Frontal fibrosing alopecia

- Graham-Little Syndrome

#### Classic pseudopelade (Brocq)

Central centrifugal cicatricial alopecia

Alopecia mucinosa

Keratosis follicularis spinulosa decalvans

 Neutrophilic
 Folliculitis decalvans

 Dissecting cellulitis/folliculitis

 Mixed
 Folliculitis (acne) keloidalis

 Folliculitis (acne) necrotica

 Erosive pustular dermatosis

 Non specific

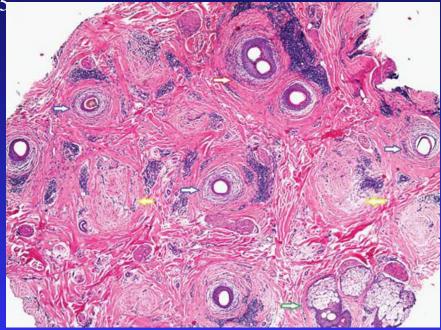
#### Central Centrifugal Cicatricial Alopecia Presenting with Irregular Patchy Alopecia on the Lateral and Posterior Scalp

Mariya Miteva Antonella Tosti

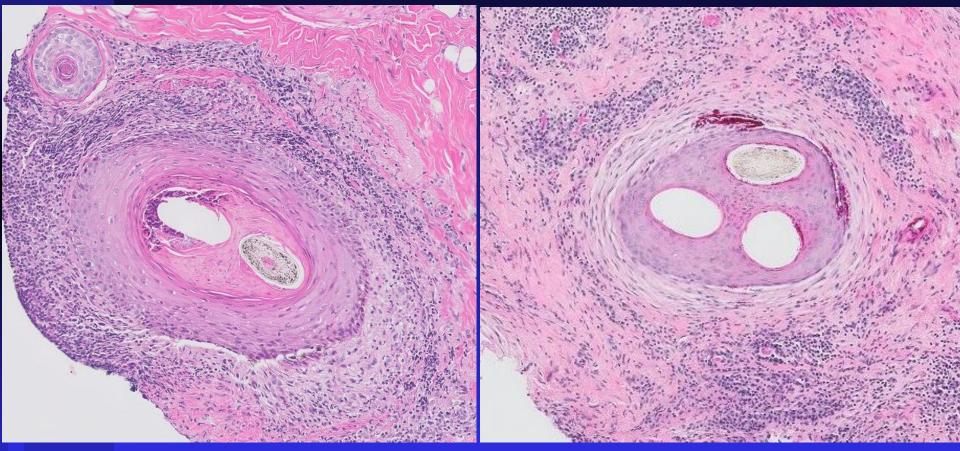
Department of Dermatology and Cutaneous Surgery, University of Miami Miller School of Medicine, Miami, Fla., USA

#### Skin Appendage Disord





### CCCA vs LPP—Current Study



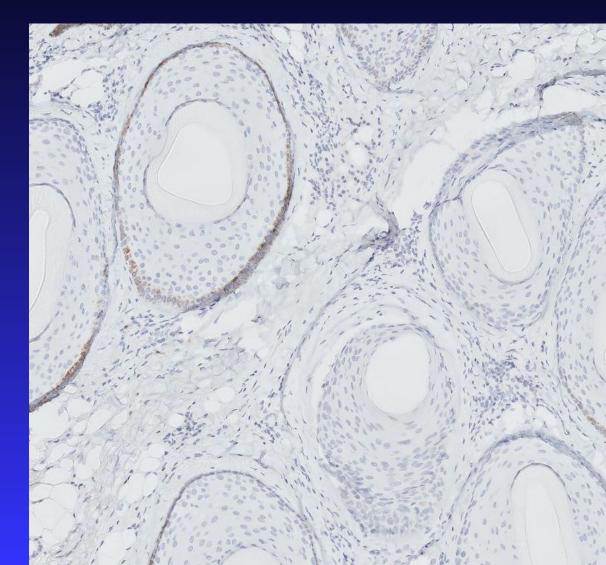




### Data collection and Analysis

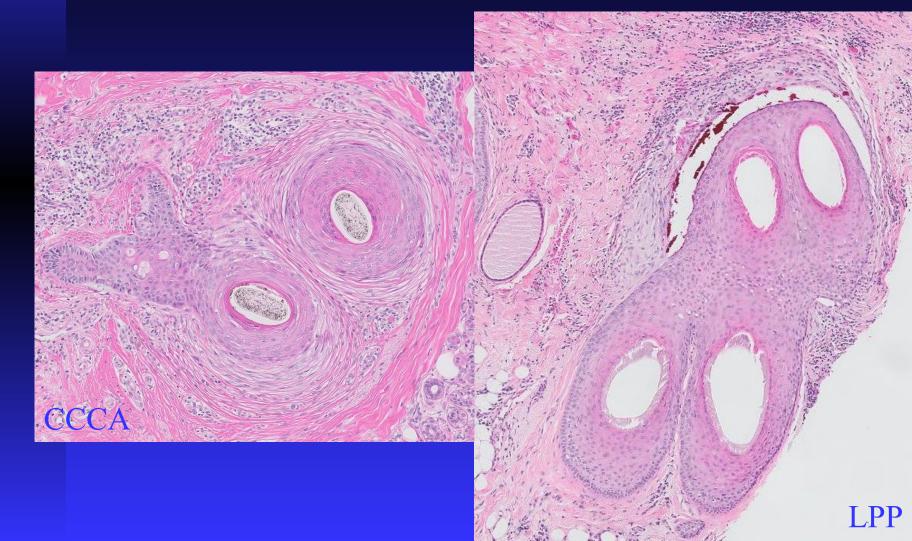
CD4, CD8, CD20, CD68,
CD123
Myeloperoxidase
CK15

### Conclusion #1 CK15 helps detects subtle disease



# Conclusion #2 Squamotization of Follicular Epithelium: Same as lichen planus Lichen planus ΙΡΡ

### Conclusion #2 Squamotization of Follicular Epithelium Identical in CCCA and LPP



Conclusion #2 Squamotization of Follicular Epithelium is "Premature Desquamation of the Inner Root Sheath"

#### Squamotization

#### Normal



### Conclusion #3 Lack of neutrophils (MPO) suggests no relationship between Folliculitis Decalvans and CCCA

SIMPLIFIED CLASSIFICATION OF PRIMARY CICATRICIAL ALOPECIA

Central Centrifugal cicatricial alopecia

Lichen planopilaris

Folliculitis decalvans neutrophilic variant of CCCA?

Sperling LC, Cowper SE, Knopp EA. An atlas of hair pathology with clinical correlations. 2nd ed. Boca Raton (FL): Taylor and Francis Group; 2012.

### Overall Conclusion: For diagnostic purposes, CCCA is identical to LPP



### Pseudopelade of Brocq



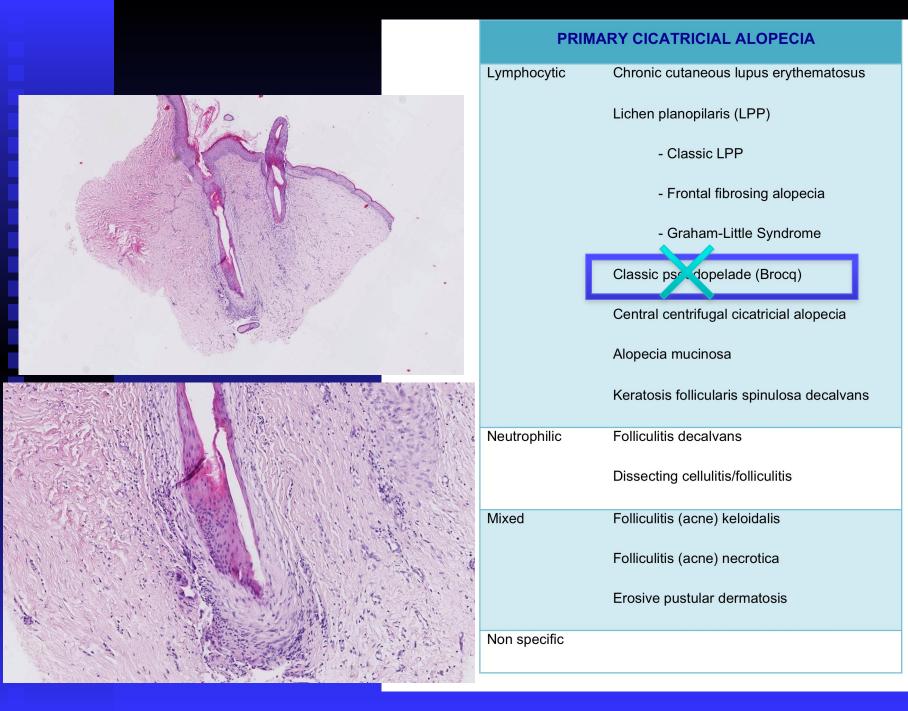
### Pseudopelade of Brocq

- With transverse sections foci of inflammation and scarring are identified
- Now diagnosed as lichen planopilaris
- Recommend abandoning this term.

### Brocq pseudopelade

## Brocq pseudopelade lichen planopilari

Silvers DN, Katz BE, Young AW. Pseudopelade of Brocq is lichen planopilaris: report of four cases that support this nosology. Cutis 1993; 51: 99.



### Objective #4

New Algorithm

North American Hair Research Society (NAHRS) Classification

#### PRIMARY CICATRICIAL ALOPECIA Chronic cutaneous lupus erythematosus Lymphocytic Lichen planopilaris (LPP) - Classic LPP - Frontal fibrosing alopecia - Graham-Little Syndrome Classic pseudopelade (Brocg) Central centrifugal cicatricial alopecia Alopecia mucinosa Keratosis follicularis spinulosa decalvans Neutrophilic Folliculitis decalvans Dissecting cellulitis/folliculitis Folliculitis (acne) keloidalis Mixed Folliculitis (acne) necrotica Erosive pustular dermatosis Non specific

- Olsen EA, Bergfeld WF, Cotsarelis G, et al. Summary of North American Hair Research Society (NAHRS)-sponsored Workshop on Cicatricial Alopecia, Duke University Medical Center, February 10 and 11, 2001. J Am Acad Dermatol 2003; 48: 103.
- 2. Olsen EA, Stenn K, Bergfeld W, et al. Update on Cicatricial Alopecia. J Invest Dermatol Symp Proc 2003; 8: 18.

## American Hair **Research Society** (AHRS) Classification

#### PRIMARY CICATRICIAL ALOPECIA

_ymphocytic	Chronic cutaneous lupus erythematosus
	Lichen planopilaris (LPP)
	- Classic LPP
	- Frontal fibrosing alopecia
	- Graham-Little Syndrome
	Classic pseudopelade (Brocq)
	Central centrifugal cicatricial alopecia
	Alopecia mucinosa
	Keratosis follicularis spinulosa decalvans
Neutrophilic	Folliculitis decalvans
	Dissecting cellulitis/folliculitis
Mixed	Folliculitis (acne) keloidalis
	Folliculitis (acne) necrotica
	Erosive pustular dermatosis



#### **PRIMARY CICATRICIAL ALOPECIA**

Lymphocytic Chronic cutaneous lupus erythematosus

Lichen planopilaris (LPP)

- Classic LPP

- Frontal fibrosing alopecia

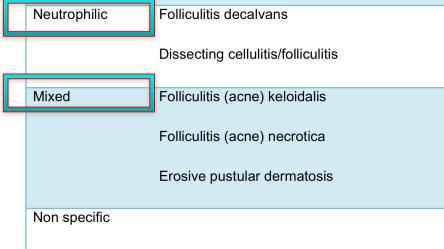
- Graham-Little Syndrome

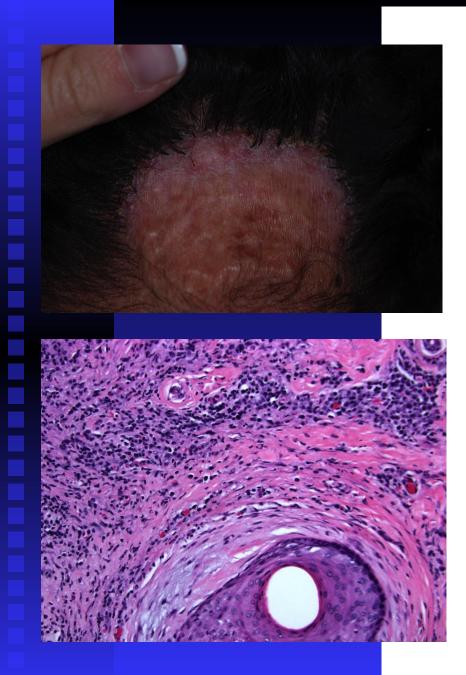
Classic pseudopelade (Brocq)

Central centrifugal cicatricial alopecia

Alopecia mucinosa

Keratosis follicularis spinulosa decalvans





#### **PRIMARY CICATRICIAL ALOPECIA**

Lymphocytic Chronic cutaneous lupus erythematosus

Lichen planopilaris (LPP)

- Classic LPP

- Frontal fibrosing alopecia

- Graham-Little Syndrome

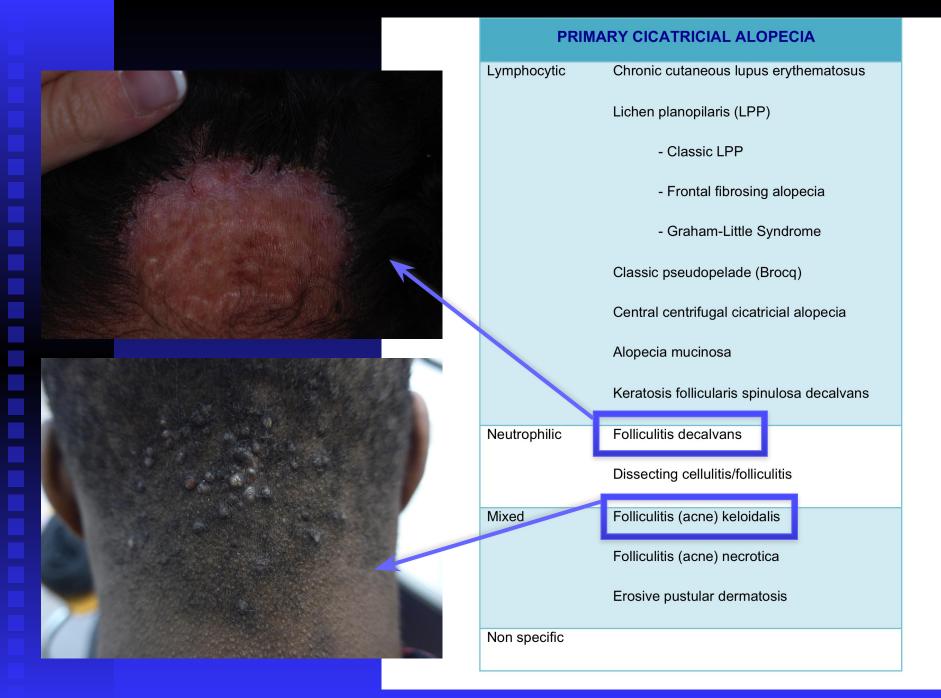
Classic pseudopelade (Brocq)

Central centrifugal cicatricial alopecia

Alopecia mucinosa

Keratosis follicularis spinulosa decalvans

Neutrophilic	Folliculitis decalvans	
	Dissecting cellulitis/folliculitis	
Mixed	Folliculitis (acne) keloidalis	
	Folliculitis (acne) necrotica	
	Erosive pustular dermatosis	
Non specific		



## Simplified classification lumps clinically disparate disorders

SIMPLIFIED CLASSIFICATION OF PRIMARY CICATRICIAL ALOPECIA Central Centrifugal cicatricial alopecia  $\neq$ Folliculitis Decalvans

Lichen planopilaris

Chronic cutaneous lupus erythematosus (discoid lupus erythematosus)

Folliculitis Keloidalis (folliculitis nuchae, acne keloidalis nuchae)

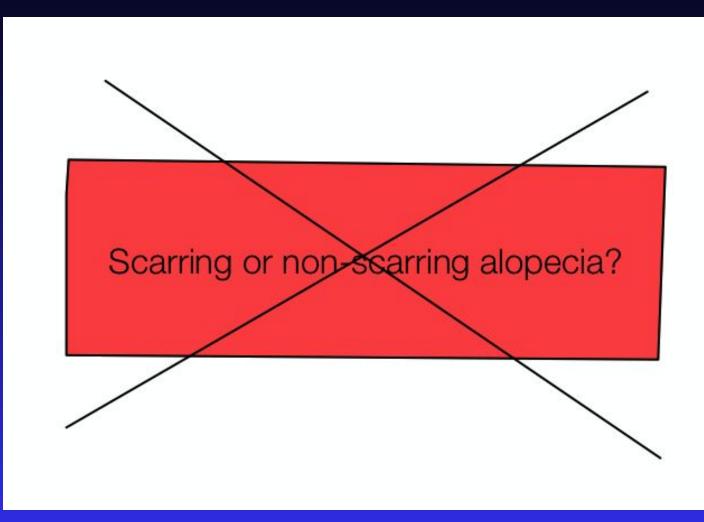
Dissecting cellulitis (perifolliculitis abscedens et suffodiens)

Cicatricial alopecia, not otherwise classified

Sperling LC, Cowper SE, Knopp EA. An atlas of hair pathology with clinical correlations. 2nd ed. Boca Raton (FL): Taylor and Francis Group; 2012.

### LIMITATIONS

- 1. All alopecias may, in the end, be scarring
- 2. Cases in marked follicular miniaturization or increase of the catagen/telogen count may be misinterpreted as cicatricial alopecia (empty tracts).
- 3. Using the cell type of the infiltrate for classification may be misleading.



## How?

How to bridge the historical classification of alopecia with all the current concepts and how to provide clinicians and dermatopathologists a practical guide to diagnosis? J Cutan Pathol 2016 doi: 10.1111/cup.12822 John Wiley & Sons. Printed in Singapore © 2016 John Wiley & Sons A/S. Published by John Wiley & Sons Ltd

Journal of Cutaneous Pathology

#### Review

# Primary scalp alopecia: new histopathological tools, new concepts and a practical guide to diagnosis

The diagnosis of primary scalp alopecia remains one of the most challenging fields in dermatopathology. In this review, we would like to connect the established classification of primary alopecia into scarring (cicatricial) and non-scarring (non-cicatricial) with current concepts. We introduce a simplified pathway for the diagnosis of the most common causes of alopecia, including a discussion of tissue processing techniques and use of immunohistochemistry.

*Keywords:* non-scarring (non-cicatricial ) alopecia, primary scalp alopecia, scarring (cicatricial) alopecia

Kolivras A, Thompson C. Primary scalp alopecia: new histopathological tools, new concepts and a practical guide to diagnosis.

J Cutan Pathol 2016. © 2016 John Wiley & Sons A/S. Published by John Wiley & Sons Ltd

#### Athanassios Kolivras<sup>1,2</sup> and Curtis Thompson<sup>3,4,5</sup>

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 <sup>2</sup>Department of Dermatopathology, Saint-Pierre, Brugmann and Queen Fabiola Children's University Hospitals, Université Libre de Bruxelles, Brussels, Belgium,
 <sup>3</sup>Department of Biomedical Engineering, Oregon Health Sciences University, Portland, OR, USA,
 <sup>4</sup>Department of Pathology, Oregon Health Sciences University, Portland, OR, USA, and

Sciences University, Portland, OR, USA, and <sup>5</sup>Department of Dermatology, Oregon Health Sciences University, Portland, OR, USA

Athanassios Kolivras, MD, PhD Departments of Dermatology and Dermatopathology, Saint-Pierre, Brugmann and Queen Fabiola Children's University Hospitals, Université Libre de Bruxelles, 129 boulevard de Waterloo, 1000 Brussels, Belgium Tel: +32 2 5354379 Fax: +32 2 5354381 e-mail: kolivras@gmail.com

Accepted for publication September 20, 2016

### A practical guide to diagnosis

• Diffuse or patchy?

Getting the clinical clues

• Usual or special clinical context?

- Examine the biopsy specimen
- Density
- Follicular size
- Catagen and telogen percentage
- Perifollicular scarring
- Inflammatory infiltrate
- Follicular fibrous tracts

Apply the 2step algorithm

- Presence or absence of follicular miniaturization?
- Increased or normal CT count?

### **Diffuse or patchy? Usual or special clinical context?**

Table 1

CLINICAL CONTEXT	DIFFUSE ALOPECIA	PATCHYALOPECIA
COMMON ENTITIES	ALOPECIA AREATA	ACNE KELOIDALIS
	FIBROSING ALOPECIA IN A PATTERN DISTRIBUTION	ALOPECIA AREATA
	LICHEN PLANOPILARIS	CENTRAL CENTRIFUGAL SCARRING ALOPECIA
	PATTERN HAIR LOSS (FEMALE AND MALE)	DISSECTING CELLULITIS
	TELOGEN EFFLUVIUM (ACUTE AND CHRONIC)	FOLLICULITIS DE CALVANS
		LICHEN PLANOPILARIS
		LUPUS ERYTHEMATOSUS (DISCOID AND NON-SCARRING)
		SYPHILIS
		TRACTION ALOPECIA (EARLY AND CHRONIC)
		TRICHOTILLOMANIA
SPECIAL ENTITIES	CHEMOTHERAPY-RELATED ALOPECIA	FRONTAL FIBROSING ALOPECIA
	I. AN AGEN EFFLUMUM	LOOSE ANAGEN SYNDROME
	II. PERMANENT ALOPECIA AFTER CHEMOTHERAPY	POST-OPERATIVE (PRESSURE-INDUCED) ALOPECIA
	LOOSE ANAGEN SYNDROME	PSORIATIC ALOPECIA AND DRUG-INDUCED PSORIASIFORM ALOPECIA
	SHORT ANAGEN SYNDROME	TRIANGULAR ALOPECIA

### **Diffuse or patchy? Usual or special clinical context?**

Table 1

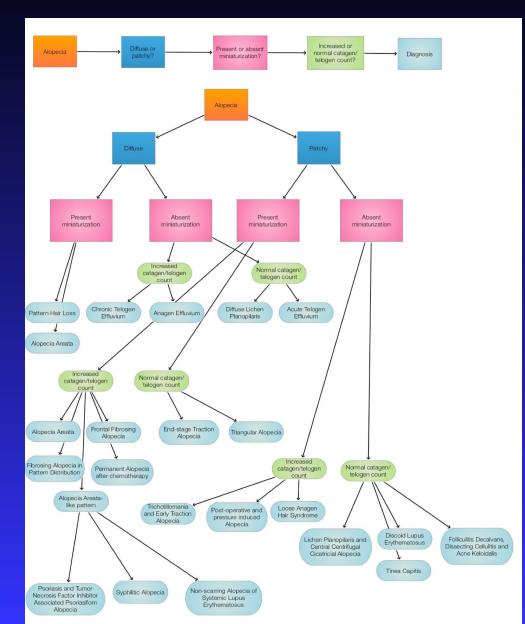
CLINICAL CONTEXT	DIFFUSE ALOPECIA	PA TCHY A LOPECIA
COMMON ENTITIES	ALOPECIA AREATA	ACNE KELOIDALIS
	FIBROSING ALOPECIA IN A PATTERN DISTRIBUTION	ALOPECIA AREATA
	LICHEN PLANOPILARIS	CENTRAL CENTRIFUGAL SCARRING ALOPECIA
	PATTERN HAIR LOSS (FEMALE AND MALE)	DISSECTING CELLULITIS
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		LICHEN PLANOPILARIS
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	SHORT ANAGEN SYNDROME	TRIANGULAR ALOPECIA

### **Diffuse or patchy? Usual or special clinical context?**

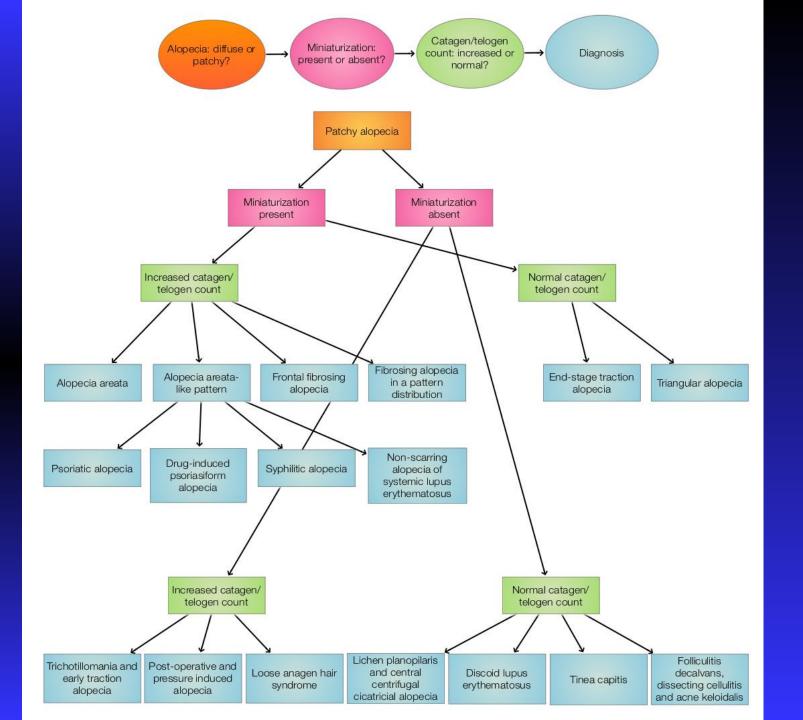
Table 1

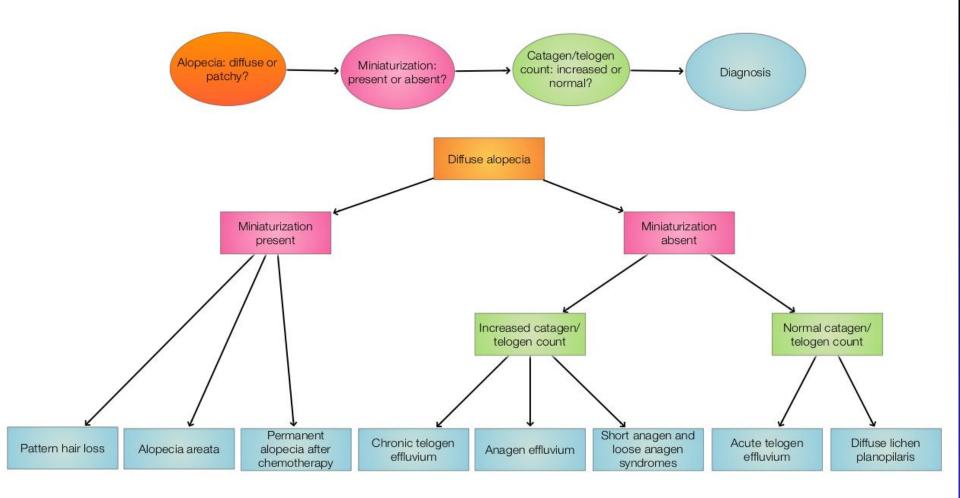
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	LOOSE ANAGEN SYNDROME	PSORIATIC ALOPECIA AND DRUG-INDUCED PSORIASIFORM ALOPECIA
	SHORT ANAGEN SYNDROME	TRIANGULAR ALOPECIA

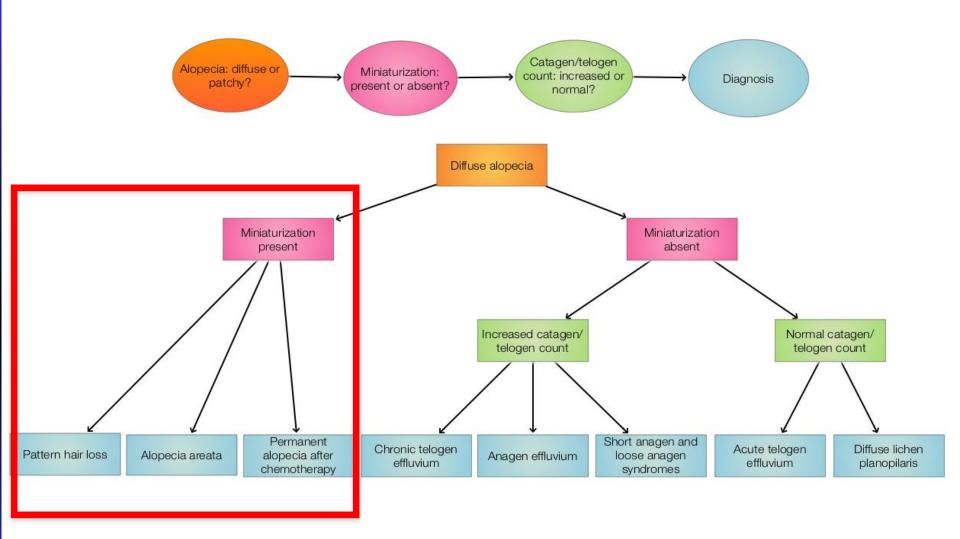
### Diagnostic Algorithm

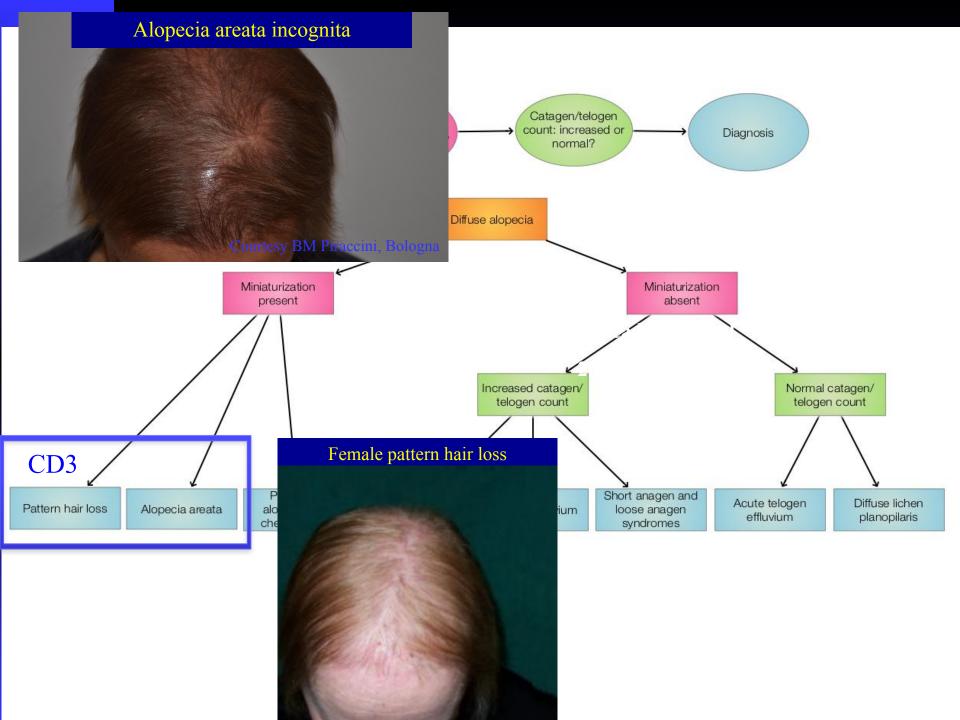


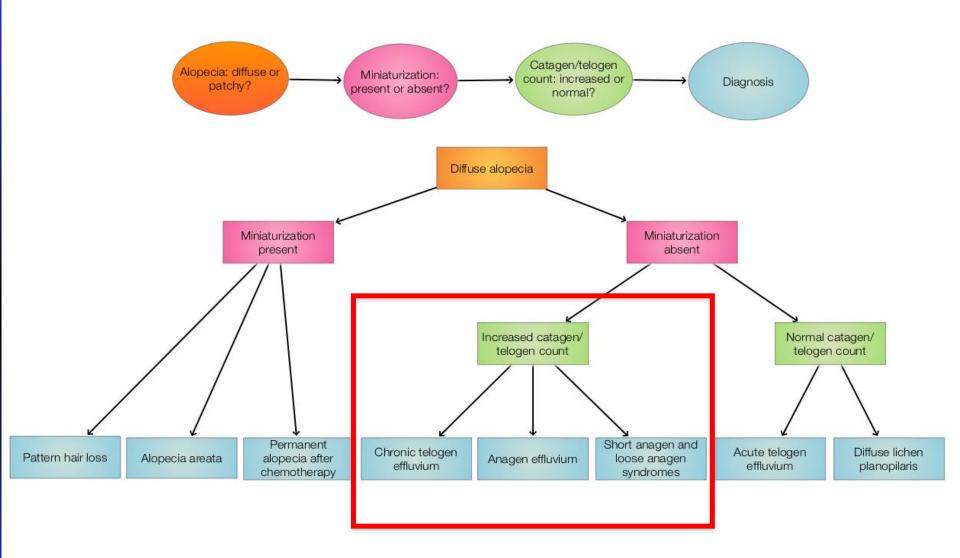


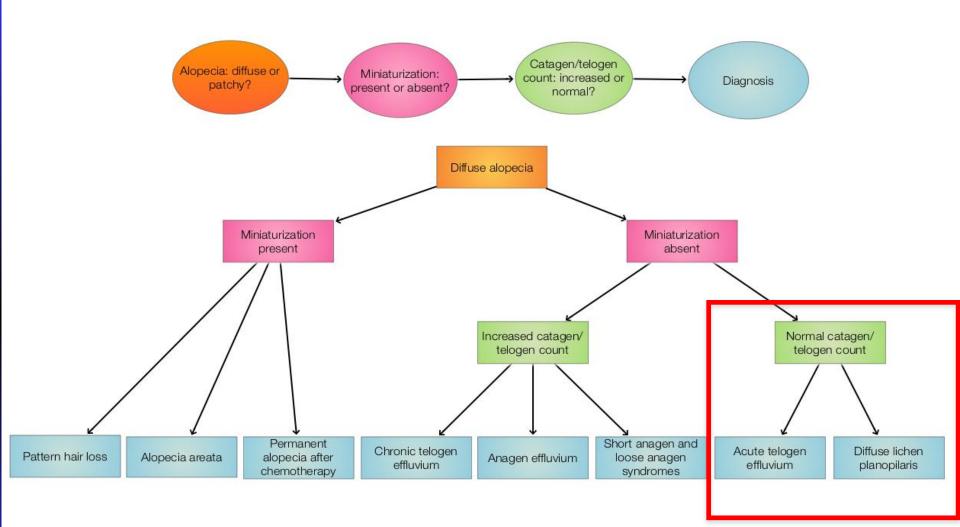




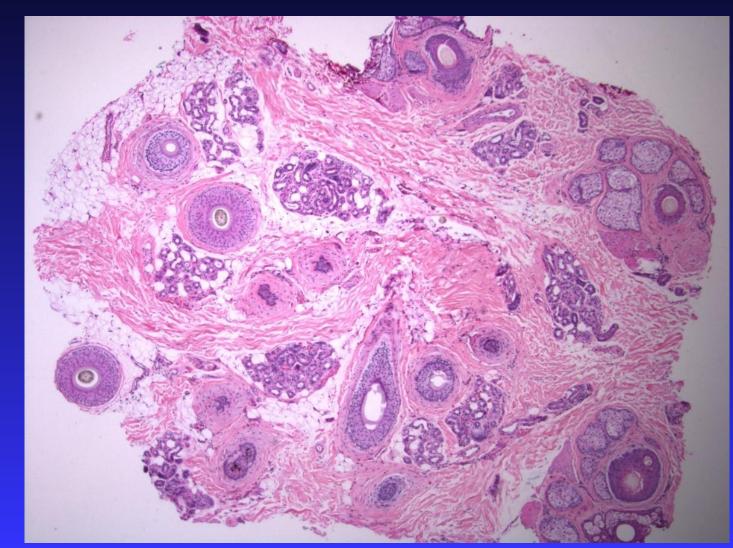


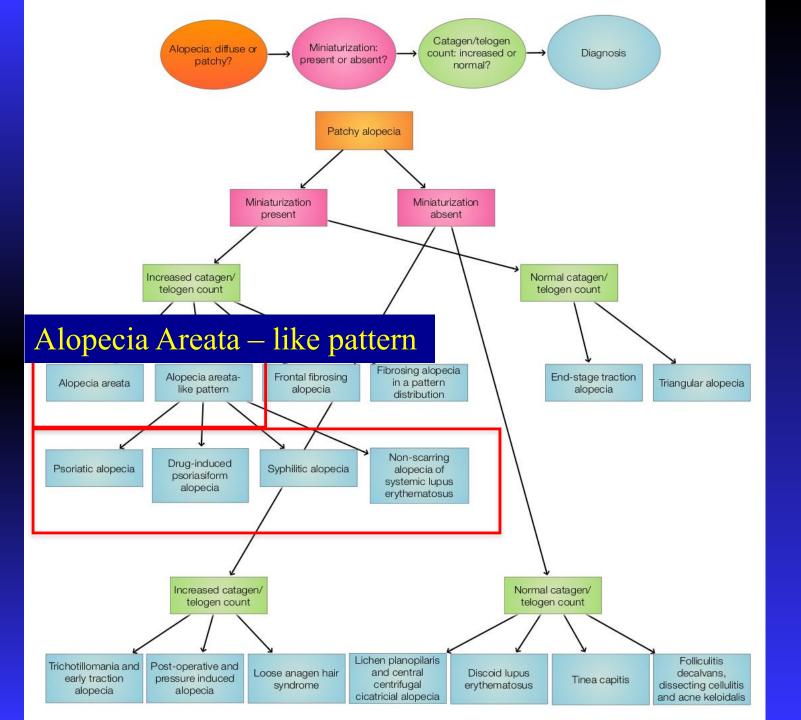






### Alopecia Areata-like Pattern can be a few of entities



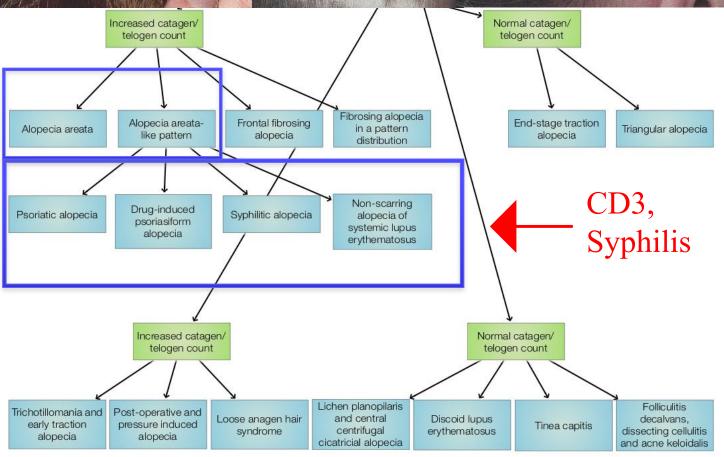


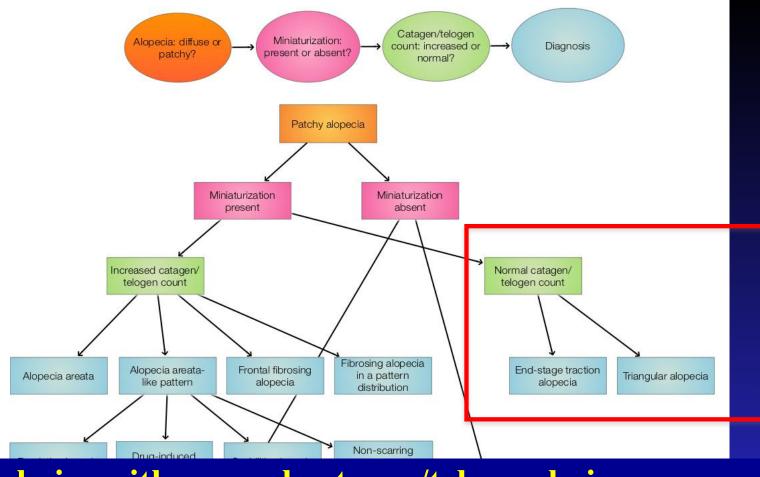
### Post-etanercept alopecia

### Syphilitic alopecia

## Non-scarring alopecia of SLE

Sperling LC, Cowper SE, Knopp EA. An Atlas of Hair Pathology with Clinical Correlations. 2<sup>nd</sup> ed. Boca Raton (FL): Taylor and Francis Group; 2012.





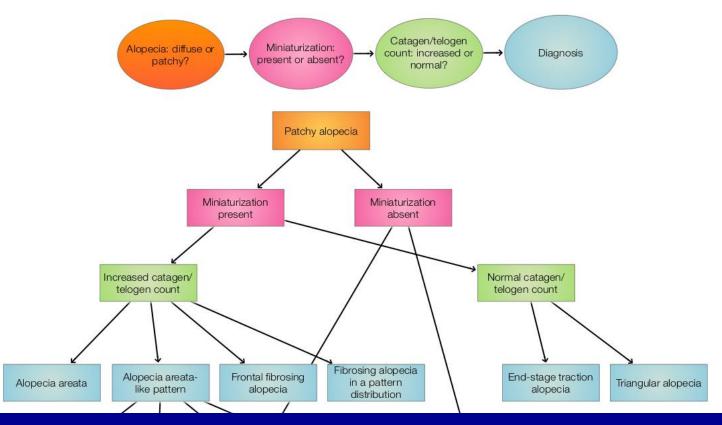
Vellus hairs with normal catagen/telogen hairs: Search for fibrous tracts 1.Triangular alopecia: absent fibrous tracts 2.Miniaturization in chronic (end-stage) traction alopecia

> Trichotillomania and Po early traction pr alopecia

Post-operative and pressure induced alopecia Loose anagen hair syndrome Lichen planopilaris and central centrifugal cicatricial alopecia

Tinea capitis

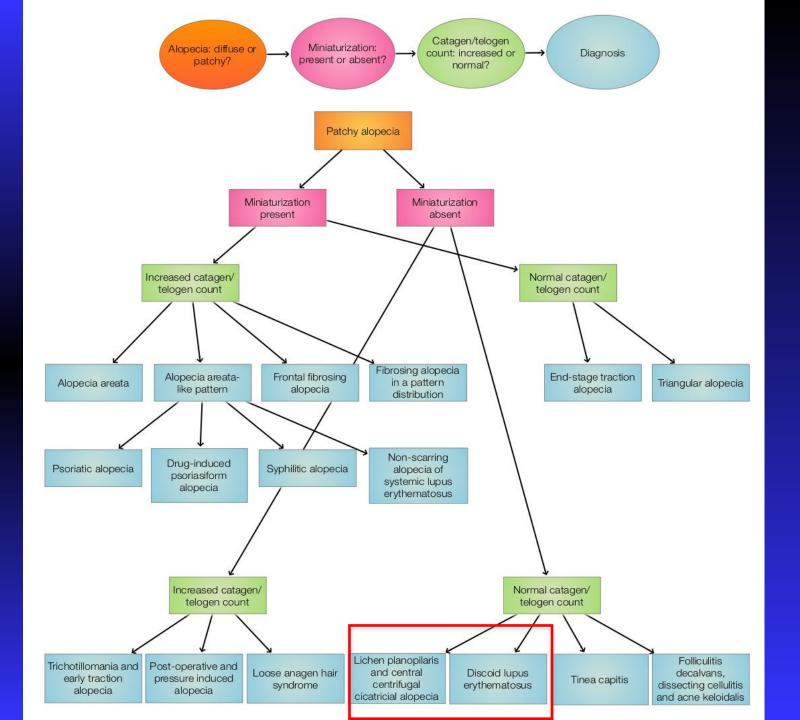
Folliculitis decalvans, dissecting cellulitis and acne keloidalis

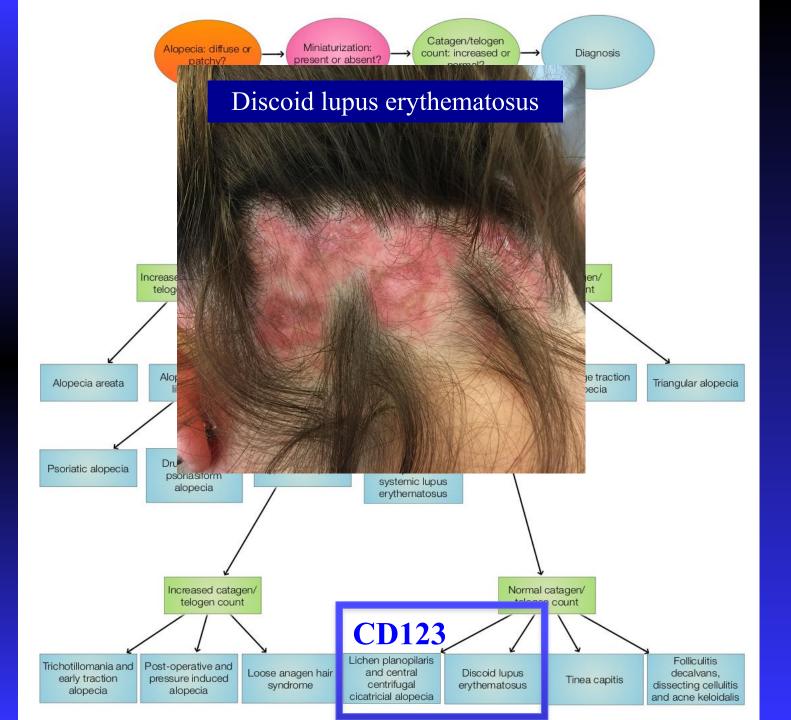


Catagen/telogen shift without follicular miniaturization
1.Trichotillosis (trichotillomania and early traction alopecia)
2.Post-operative and pressure induced alopecia: vascular thrombosis and fat necrosis

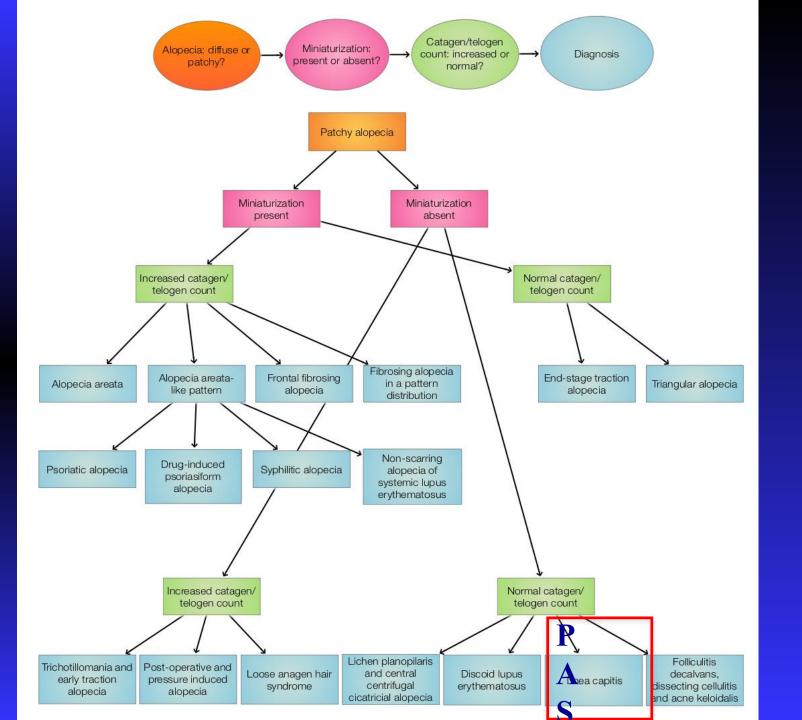




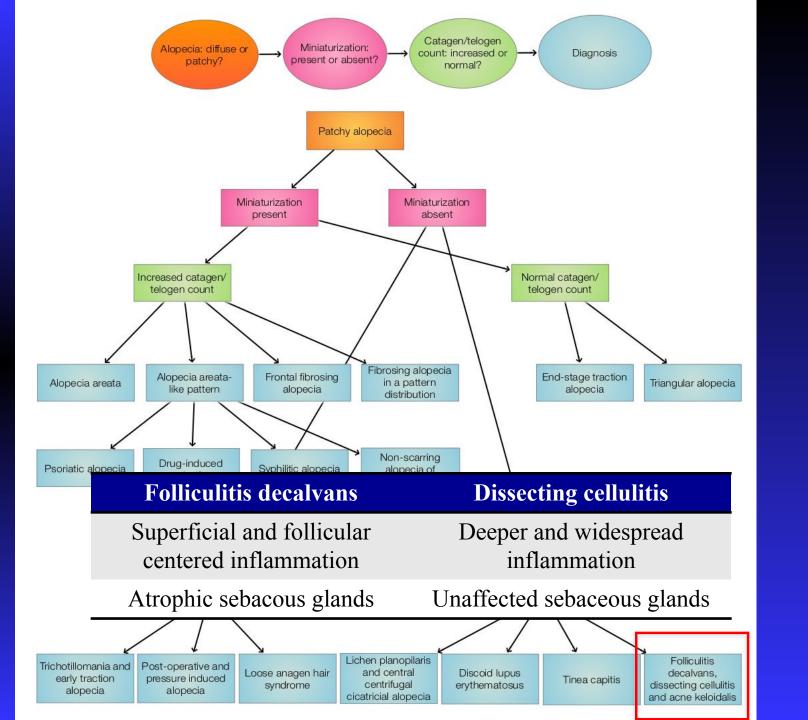




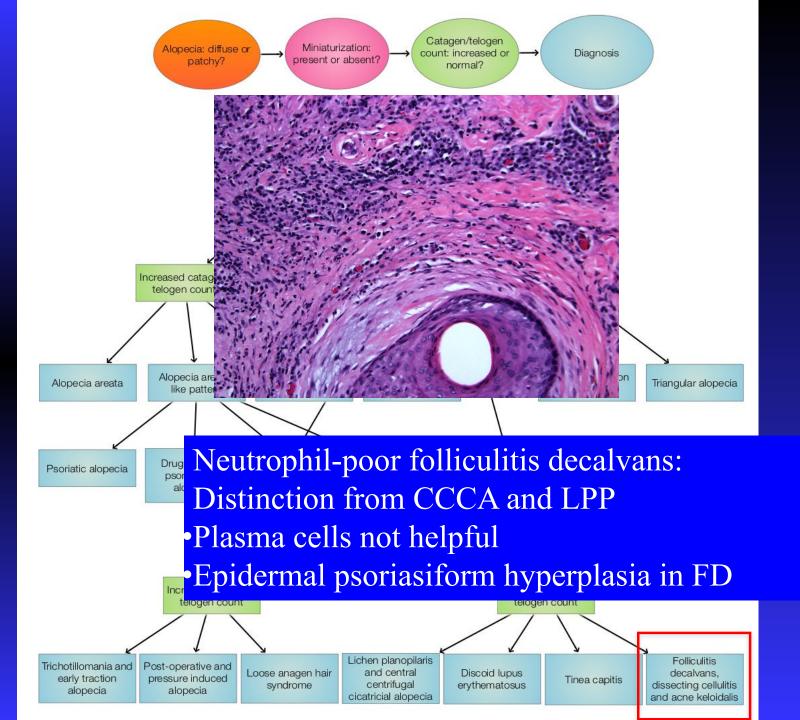


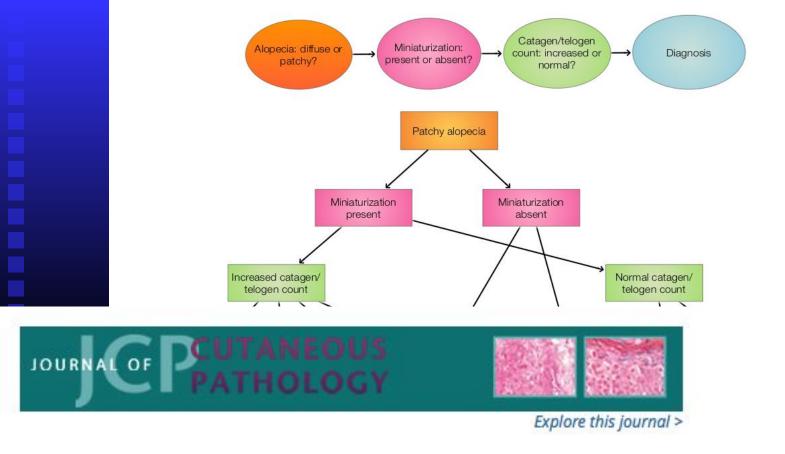






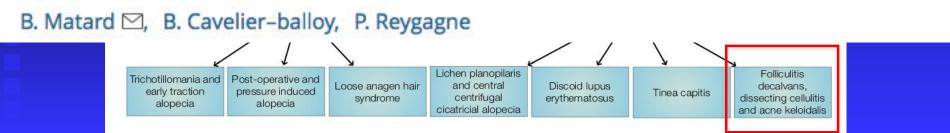






**ORIGINAL ARTICLE** 

## Epidermal psoriasiform hyperplasia, an unrecognized sign of folliculitis decalvans: a histological study of 26 patients

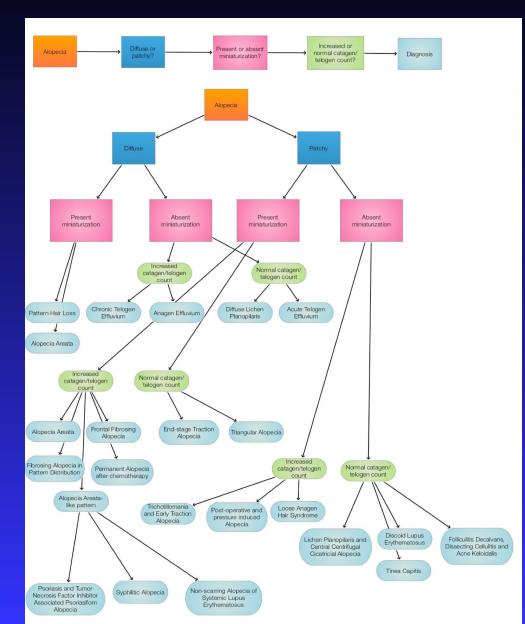


## **Diffuse or patchy? Usual or special clinical context?**

Table 1

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	SHORT ANAGEN SYNDROME	TRIANGULAR ALOPECIA

# Diagnostic Algorithm



## Acknowledgements

- Athanassios Kolivras, MD, Saint-Pierre, Brugmann and HUDERF Hospitals, Université Libre de Bruxelles, Brussels, Belgium.
- Janet Roberts, MD and Nisha Desai, MD
- **Residents and Medical Students** 
  - Cindi Chapman, MD
  - Arlette Habashi-Daniel, MD
  - Elizabeth Dawson, MD (Caroline Xavier, MD) and Omar Lupi, MD-Rio de Janeiro)
- Vera Price, MD







# Thank you!

## MAY 17TH, 2017

### UPDATED 7:00 AM



**28°** 

Snowing

### **TODAY AT TIMBERLINE:**

We are scheduled to operate Today

NEW SNOW	INCHES
SINCE 5AM:	1"
LAST 24 HRS (5am to 5am):	6"
LAST 72 HRS:	12"
BASE DEPTH AT LODGE:	175"
Click for Palmer Snow Depth	



Please come visit!

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