#### Azienda Ospedaliera - Università di Padova Centro Regionale di Dermatologia Pediatrica

## Eczema e Psoriasi, qui va sempre peggio!

Anna Belloni Fortina





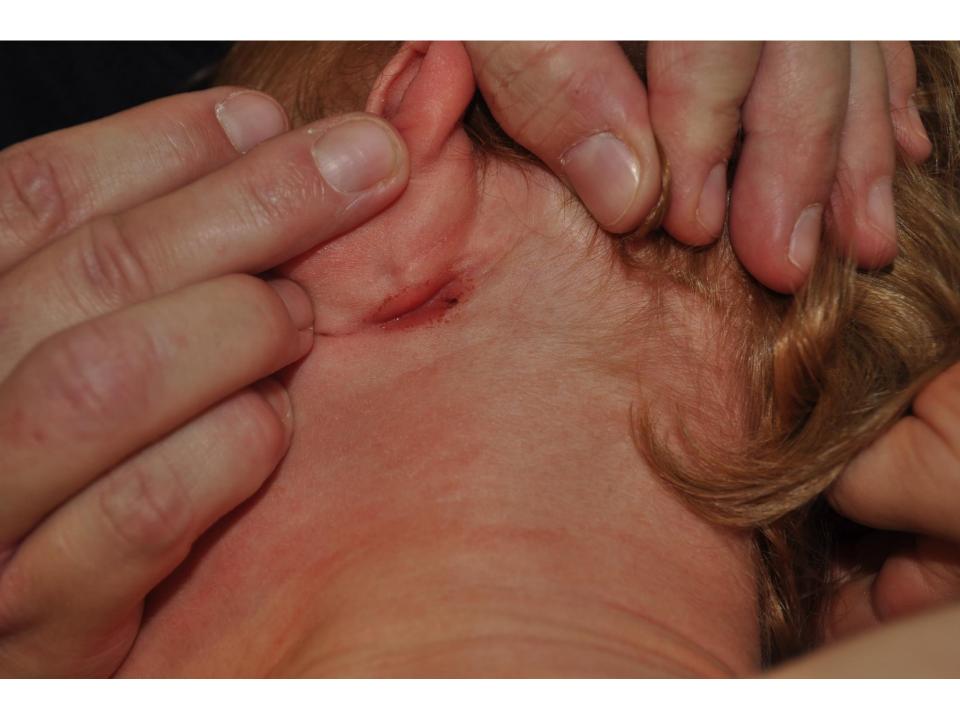




















# Eczema atopico o Psoriasi?

**Psoriasi** 











Review

### Diagnosis of Atopic Dermatitis: Mimics, Overlaps, and Complications

Elaine C. Siegfried 1,†,\* and Adelaide A. Hebert 2,†

	Diagnosis	Relative Prevalence			
	Diagnosis	Infants	Children	Adolescents/Adults	
	seborrheic dermatitis	common	uncommon	common	
	psoriasis	less common	less common	common	
Inflammatory Skin	nummular dermatitis	less common	common	less common	
<b>Conditions</b>	contact dermatitis <sup>a</sup>	common	common	common	
(See Section 2)	dermatographism <sup>a</sup>	less common	common	common	
	pityriasis alba <sup>a</sup>	common	common	uncommon	
	overlap (see Section 2.7)	common	common	common	

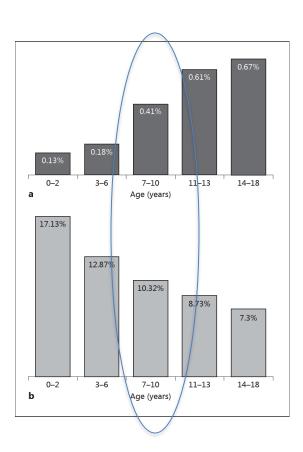
"Overlap" is a term used to describe one or more coexisting inflammatory skin diseases. The most well described combination may be psoriasis-eczema overlap (PsE), also known as eczematous psoriasis and PsEma. Patients with PsE typically present with a combination of flexural eczema and psoriatic lesions that lack thick plaques and are more likely to experience itch than patients with isolated psoriasis (Figure 8) [12,34,35]. In one study, PsE responded well to psoriasis treatment strategies including TCS [34].

#### **Dermatology**

Dermatology 2015;231:35–40 DOI: 10.1159/000381913 Received: March 3, 2015 Accepted: March 26, 2015 Published online: May 8, 2015

## **Epidemiology and Comorbidity in Children with Psoriasis and Atopic Eczema**

Matthias Augustin<sup>a</sup> Marc A. Radtke<sup>a</sup> Gerd Glaeske<sup>c</sup> Kristian Reich<sup>b</sup> Enno Christophers<sup>d</sup> Ines Schaefer<sup>a</sup> Arnd Jacobi<sup>a</sup>

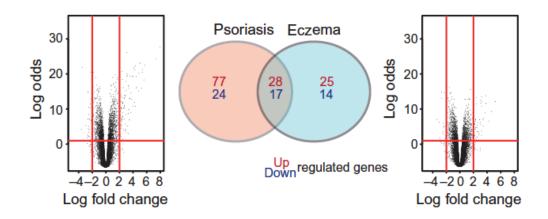


Diagnosis	With PSO	Without PSO
Arthritis	0.15	0.01
Iridocyclitis	0.38	0.04
Alopecia areata	0.46	0.12
Nail disorders	2.89	0.93
Ulcerative colitis	0.15	0.05
Atopic eczema	24.52	10.29
Impetigo	3.35	1.47
Contact dermatitis	3.66	1.73
Arterial hypertension	0.91	0.44
Vitiligo	0.23	0.11
Diabetes mellitus	0.61	0.31
Obesity	7.08	3.61
Hyperlipidaemia	1.14	0.64
Viral warts	12.11	7.29
Depression	1.29	0.77
Keratitis	0.53	0.32
Allergic rhinitis	15.16	9.70
Bronchial asthma	12.19	9.36
Herpes infection	2.74	2.12
Ischaemic heart disease	0.08	0.06
ADHD	8.07	7.14
Urticaria	2.28	2.03
Chronic bronchitis	2.21	2.10
Crohn's disease	0.00	0.07

#### **PSORIASIS**

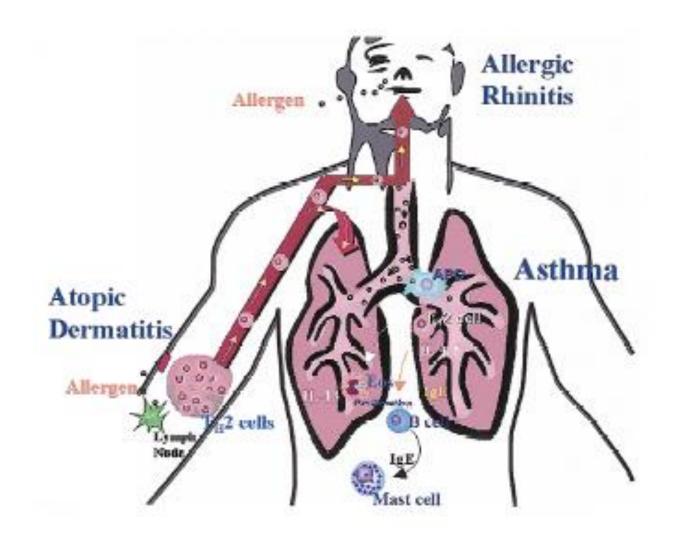
# Intraindividual genome expression analysis reveals a specific molecular signature of psoriasis and eczema

Maria Quaranta,<sup>1\*</sup> Bettina Knapp,<sup>2\*</sup> Natalie Garzorz,<sup>3</sup> Martina Mattii,<sup>1</sup> Venu Pullabhatla,<sup>4</sup> Davide Pennino,<sup>1</sup> Christian Andres,<sup>3</sup> Claudia Traidl-Hoffmann,<sup>3</sup> Andrea Cavani,<sup>5</sup> Fabian J. Theis,<sup>2,6</sup> Johannes Ring,<sup>3</sup> Carsten B. Schmidt-Weber,<sup>1</sup> Stefanie Eyerich,<sup>1†</sup> Kilian Eyerich<sup>3†‡</sup>





## "atopic march"



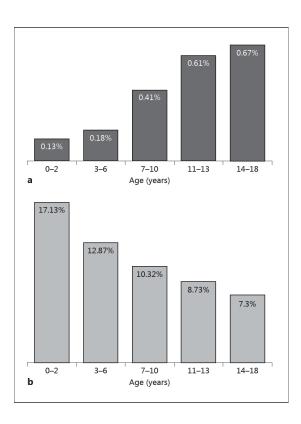
Spergel & Paller – J Allergy Clin Immunol 2003; 112:S118-S127

#### **Dermatology**

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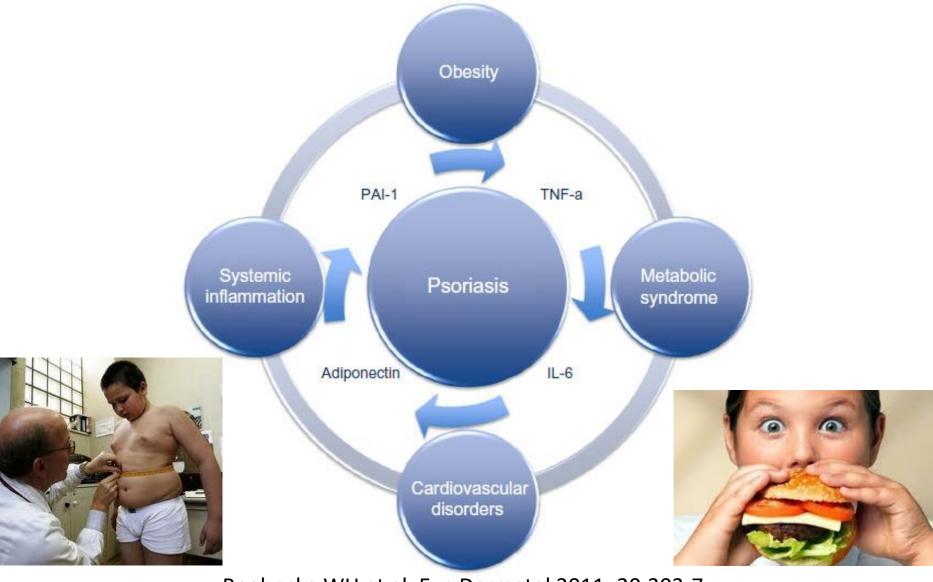
## **Epidemiology and Comorbidity in Children with Psoriasis and Atopic Eczema**

Matthias Augustin<sup>a</sup> Marc A. Radtke<sup>a</sup> Gerd Glaeske<sup>c</sup> Kristian Reich<sup>b</sup> Enno Christophers<sup>d</sup> Ines Schaefer<sup>a</sup> Arnd Jacobi<sup>a</sup>



Diagnosis	With PSO	Without PSO	With AE	Without AE
Arthritis	0.15	0.01	0.01	0.01
Iridocyclitis	0.38	0.04	0.06	0.03
Alopecia areata	0.46	0.12	0.21	0.09
Nail disorders	2.89	0.93	1.29	0.78
Ulcerative colitis	0.15	0.05	0.07	0.04
Atopic eczema	24.52	10.29	100.00	0.00
Impetigo	3.35	1.47	3.57	1.07
Contact dermatitis	3.66	1.73	3.61	1.32
Arterial hypertension	0.91	0.44	0.40	0.39
Vitiligo	0.23	0.11	0.26	0.08
Diabetes mellitus	0.61	0.31	0.34	0.26
Obesity	7.08	3.61	4.11	3.10
Hyperlipidaemia	1.14	0.64	0.71	0.55
Viral warts	12.11	7.29	9.76	6.10
Depression	1.29	0.77	0.83	0.67
Keratitis	0.53	0.32	0.33	0.28
Allergic rhinitis	15.16	9.70	19.64	7.44
Bronchial asthma	12.19	9.36	19.04	7.16
Herpes infection	2.74	2.12	3.41	1.71
Ischaemic heart disease	0.08	0.06	0.08	0.05
ADHD	8.07	7.14	8.11	6.10
Urticaria	2.28	2.03	3.61	1.60
Chronic bronchitis	2.21	2.10	3.88	1.64
Crohn's disease	0.00	0.07	0.08	0.06

### "psoriatic march"



Boehncke WH et al. Exp Dermatol 2011; 20:303-7

Tiago TORRES<sup>1,2</sup> Susana MACHADO<sup>1,3</sup> Denisa MENDONÇA<sup>4</sup> Manuela SELORES<sup>1,3</sup>

## Cardiovascular comorbidities in childhood psoriasis

matory diseases. Results: Psoriatic children had a significantly higher prevalence and greater odds of excess adiposity compared to controls: BMI (≥85th percentile; OR 4.4; 95%CI 1.2-15.6), waist circumference (>75th percentile; OR 7.4; 95%CI 2.0-27.7) and waist-to-height ratio (>0.490; OR 4.6; 95%CI 1.3-17.0). A higher prevalence of metabolic syndrome was observed in children with psoriasis compared to controls (25% vs 3.7%; P = 0.07), and two components of the metabolic syndrome were significantly higher in the psoriasis group: waist circumference (75% vs 29.6%; P = 0.002) and the high blood pressure component (30% vs 3.7% P = 0.032). Finally, an altered and more atherogenic lipid profile was observed among psoriatic patients without excess adiposity. Conclusion: This study demonstrates that comorbidities known to be associated with adult psoriasis are also observed in childhood psoriasis, reinforcing the need for screening cardiovascular comorbidities in children with psoriasis and promoting healthy lifestyle choices in these patients. Moreover, it also suggests that its association with psoriasis may be in part genetically determined rather than uniquely acquired.



### Association of Pediatric Psoriasis Severity With Excess and Central Adiposity. An International Cross-Sectional Study

Amy S. Paller, MD; Katherine Mercy, MD; Mary J. Kwasny, ScD; Siew Eng Choon, FRCP; Kelly M. Cordoro, MD; Giampiero Girolomoni, MD; Alan Menter, MD; Wynnis L. Tom, MD; Anne M. Mahoney, MD; Annet M. Oostveen, MD; Marieke M. B. Seyger, MD

*JAMA Dermatol.* 2013;149(2):166-176.

**Objective** To investigate the relationship of excess and central adiposity with pediatric psoriasis severity.

**Design, Setting, and Participants** Multicenter, cross-sectional study **of 409 psoriatic children**. Psoriasis was classified as mild (worst Physician's Global Assessment score ≤3 with body surface area ≤10%) or severe (worst Physician's Global Assessment score ≥3 with body surface area >10%). Children were enrolled from 9 countries between June 19, 2009, and December 2, 2011.

Main Outcome Measures Excess adiposity (body mass index percentile) and central adiposity (waist circumference percentile and waist to height ratio).

Results Excess adiposity (body mass index ≥85th percentile) occurred in 37.9% of psoriatic children (n = 155) vs 20.5% of controls (n = 42) but did not differ significantly by severity. The odds ratio (95% CI) of obesity (body mass index ≥95th percentile) overall in psoriatic children vs controls was 4.29 (1.96-9.39) and was higher with severe (4.92; 2.20-10.99) than with mild (3.60; 1.56-8.30) psoriasis, particularly in the United States (7.60; 2.47-23.34, and 4.72; 1.43-15.56, respectively). Waist circumference above the 90th percentile occurred in 9.3% of the control (n = 19), 14.0% of the mild psoriasis (n = 27), and 21.2% of the of severe psoriasis (n = 43) participants internationally; this incidence was highest in the United States (12.0% [n = 13], 20.8% [16], and 31.1% [32], respectively). Waist to height ratio was significantly higher in psoriatic (0.48) vs control (0.46) children but was unaffected by psoriasis severity. Children with severe psoriasis at its worst, but mild at enrollment, showed no significant difference in excess or central adiposity from children whose psoriasis remained severe.

<u>Conclusions:</u> Globally, children with psoriasis have excess adiposity and increased central adiposity regardless of psoriasis severity. The increased metabolic risks associated with excess and central adiposity warrant early monitoring and life style modification



## Risks of developing psychiatric disorders in pediatric patients with psoriasis

Alexa B. Kimball, MD,<sup>a</sup> Eric Q. Wu, PhD,<sup>b</sup> Annie Guérin, MSc,<sup>b</sup> Andrew P. Yu, PhD,<sup>b</sup> Magda Tsaneva, BA,<sup>b</sup> Shiraz R. Gupta, PharmD,<sup>c</sup> Yanjun Bao, PhD,<sup>c</sup> and Parvez M. Mulani, PhD<sup>c</sup> Boston, Massachusetts, and Abbott Park, Illinois

**Background:** Symptoms of psoriasis can be embarrassing and distressing, and may increase risk of developing psychiatric disorders in young people.

**Objective:** We sought to compare incidences of psychiatric disorders between pediatric patients with psoriasis and psoriasis-free control subjects.

**Methods:** Patients (<18 years) with continuous health plan enrollment 6 months before and after first psoriasis diagnosis (index date) were selected (Thomson Reuters MarketScan database, 2000-2006 [Thomson Reuters, New York, NY]). Patients with psoriasis (N = 7404) were matched 1:5 on age and sex to psoriasis-free control subjects (N = 37,020). Patients were followed from index date to first diagnosis of a psychiatric disorder (ie, alcohol/drug abuse, depression, anxiety disorder, bipolar disorder, suicidal ideation, eating disorder), end of data availability, or disenrollment. Patients with psychiatric diagnoses or psychotropic medication use before the index date were excluded. Cox proportional hazard models controlling for age, sex, and comorbidities were used to estimate the effect of psoriasis on risks of developing psychiatric disorders.

**Results:** Patients with psoriasis were significantly more at risk of developing psychiatric disorders versus control subjects (5.13% vs 4.07%; P = .0001; hazard ratio = 1.25; P = .0001), especially depression (3.01% vs 2.42%; P = .0036; hazard ratio = 1.25; P = .0053) and anxiety (1.81% vs 1.35%; P = .0048; hazard ratio = 1.32; P = .0045).

*Limitations:* Retrospective, observational studies of medical claims data are typically limited by overall quality and completeness of data and accuracy of coding for diagnoses and procedures.

**Conclusions:** Pediatric patients with psoriasis had an increased risk of developing psychiatric disorders, including depression and anxiety, compared with psoriasis-free control subjects. (J Am Acad Dermatol 2012;67:651-7.)

Key words: anxiety; comorbidity; depression; pediatrics; psoriasis; psychotropic medications.



Allergy

#### REVIEW ARTICLE

Allergy 70 (2015) 887-896

#### Past, present, and future for biologic intervention in atopic dermatitis

M. D. Howell, M. L. Parker, T. Mustelin & K. Ranade

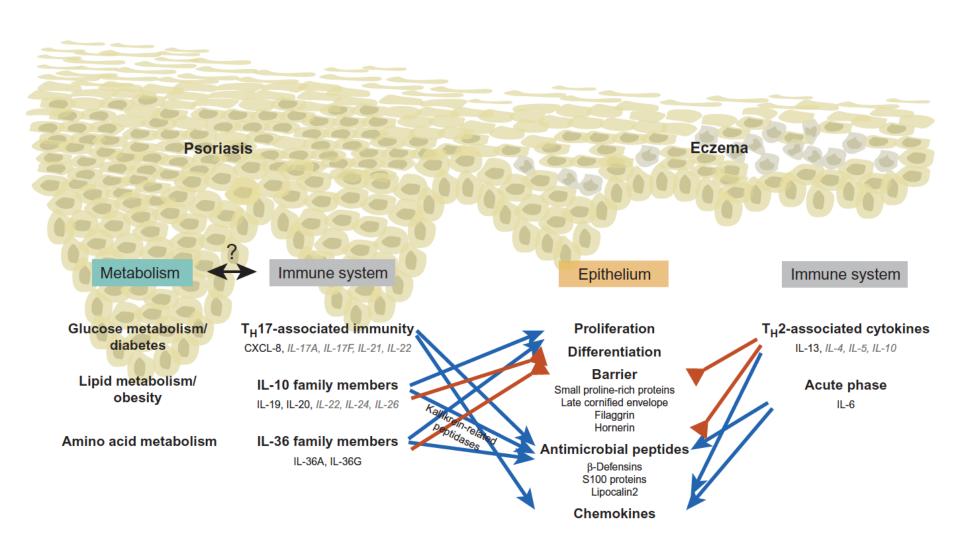
Table 2 Studies in atopic dermatitis using biologic therapies approved for other diseases

Immunologic target	Agent	Manufacturer	Approved for	References
CD2/LFA-3	Alefacept (Amevive®)	Astellas	Psoriasis	(64, 65)
CD11a/LFA-1	Efalizumab (Raptiva®)	Genentech/MerckSerono	Psoriasis*	(66, 118, 119)
CD20	Rituximab (Rituxan®)	Genentech/Biogen Idec	Chronic Lymphocytic Leukemia Non-Hodgkin's Lymphoma Rheumatoid Arthritis	(57, 58)
lgE	Omalizumab (Xolair®)	Genentech/Novartis	Asthma Chronic Idiopathic Urticaria	(52–55, 120, 121)
IL-6R	Tocilizumab (Actemra®)	Hoffman-La Roche/Genentech	Juvenile Idiopathic Arthritis Rheumatoid Arthritis	(107)
IL-12/23p40	Ustekinumab (Stelara®)	Janssen	Psoriasis Psoriatic Arthritis	(114, 115, 122)
TNF-alpha	Etanercept (Enbrel®)	Amgen	Ankylosing Spondylitis Psoriasis Psoriatic Arthritis Rheumatoid Arthritis	(104, 123)
	Infliximab (Remicade®)	Janssen	Ankylosing Spondylitis Crohn's Disease Psoriasis Psoriatic Arthritis Rheumatoid Arthritis	(103)

<sup>\*</sup>Removed from the market due to risk of developing progressive multifocal leukoencephalopathy (PML).

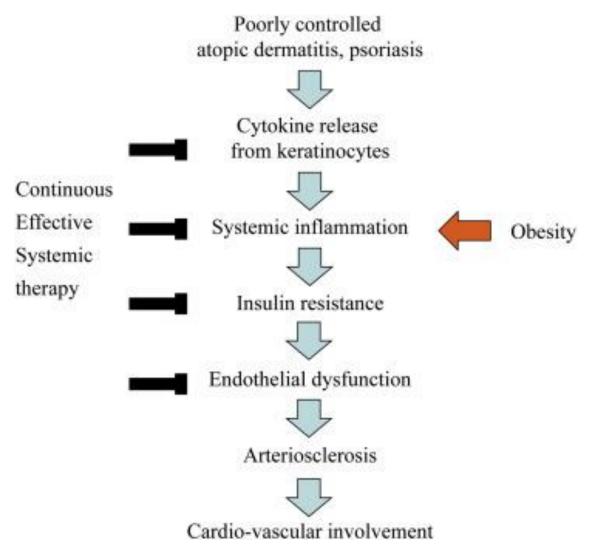
### Biologics for AD and Pso

Agent	Target	Route	Drug	Phase AD	Phase Pso	ClinicalTrials.gov
Dupilumab	II-4Ra	SC	mAb	III		NCT01949311
Ustekinumab	IL-12/23p40	SC	mAb	II	Approved	NCT01806662
Secukinumab	IL-17A	SC	mAb		Approved	
Adalimumab	Anti TNFalfa	SC	mAb		Approved	
Infliximab and biosimilar	Anti TNFalfa	EV	mAb		Approved	
Etanercept	Anti TNFalfa	SC	Fusion protein		Approved	
Apremilast	PDE4	Oral	Small molecule	Ш	Approved	NCT02087943
Tofacitinib	JAK	Topical	Small molecule	II	II	
Acitertina	Retinoid Acid receptor	Oral	Small molecule		Approved	
Tazarotene	Retinoid Acid receptor	Topical	Small molecule		Approved	





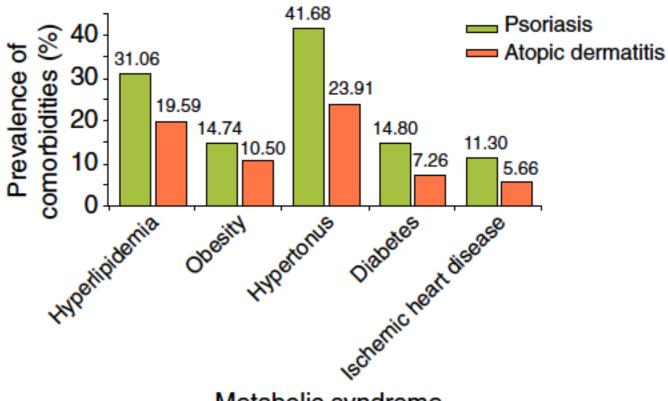
#### "Inflammatory skin march"



Keiichi Yamanaka, Hitoshi Mizutani

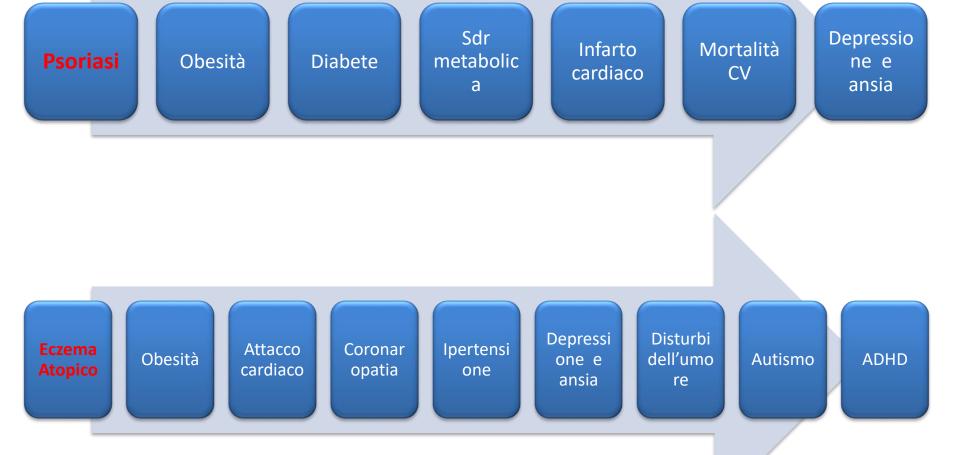
"Inflammatory skin march": IL-1-mediated skin inflammation, atopic dermatitis, and psoriasis to cardiovascular events

Journal of Allergy and Clinical Immunology, 136, 3, 2015, 823–824



Metabolic syndrome

### "Inflammatory skin march"



### Take home message

- L'eczema e la psoriasi possono coesistere, anche se in una piccola parte dei soggetti.
- È necessario riconoscerli e trattarli entrambi
- Le comorbidità devono essere valutate e monitorate anche in età pediatrica per garantire una vita futura soddisfacente
- Il trattamento è sempre utile anche per evitare la "marcia dell'infiammazione cronica cutanea"





#### Congresso Nazionale S.I.Der.P.

Bologna, 12-14 ottobre 2017

