- Hegemann L, Forstinger C, Partsch B *et al.* (1995) Microdialysis in cutaneous pharmacology kinetic-analysis of transdermally delivered nicotine. *J Investig Dermatol* 104:839-43
- Jynge P, Skjetne T, Gribbestad I *et al.* (1990) *In vivo* tissue pharmacokinetics by fluorine magnetic-resonance spectroscopy a study of liver and muscle disposition of fleroxacin in humans. *Clin Pharmacol Ther* 48:481–9
- Nowak A, Klimowicz A (1990) 2-Stage penetration of a single oral dose of sulfadimethoxine into skin blister fluid. *Eur J Clin Pharmacol* 39:487–90
- Pershing LK, Nelson JL, Corlett JL *et al.* (2003) Assessment of dermatopharmacokinetic approach in the bioequivalence determination of topical tretinoin gel products. *J Am Acad Dermatol* 48: 740–51
- Roos K, Brorson JE (1990) oncentration of phenoxymethylpenicillin in tonsillar tissue. *Eur J Clin Pharmacol* 39:417–8
- Weigmann HJ, Lademann J, Von Pelchrzim R et al. (1999) Bioavailability of clobetasol propionate—quantification of drug concentrations in the stratum corneum by dermatopharmacokinetics using tape stripping. *Skin Pharmacol Appl Skin Physiol* 12: 46-53

## Quality of Life in Alopecia Areata: A Study of 60 Cases

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#### **TO THE EDITOR**

Alopecia areata (AA) is a chronically relapsing skin disorder characterized by a sudden loss of hair. Because the perception of patients may differ significantly from those of their health-care providers, guality of life (Qol) appears to be a more relevant criterion to assess the severity of this disease than clinical evaluation such as AA extension. To our knowledge, only one Turkish study investigated the impact of AA on Qol using short form 36 (SF36), indicating lower Qol levels compared with sex-matched individuals (Gulec et al., 2004). In this study, Qol was assessed using a generic instrument. Because only three dimensions were affected and results may be linked to the specific culture, a confirmation was needed. We used an approach combining generic and specific measures to assess the impact of AA on French patients' Qol, to compare Qol levels with those observed in the general population and in other dermatological conditions, and finally to determine the impact of clinical characteristics and sociodemographic factors on Qol.

Subjects were aged over 16 years, presenting with a minimum of 8 weeks AA history, having given informed consent to participate, and having the French language as their native lan-

guage. Sociodemographic data and characteristics of the disease (duration and course, treatments in the recent period, affected surfaces on the scalp and other areas involved) were recorded. The severity of each AA was reported using visual analogical scales (0-10) by reference to (i) all the AA cases seen in daily practice; (ii) all cases of all skin disorders. Three selfadministered questionnaires were used to assess Qol: the generic and worldwide-used SF36 (Leplege et al., 1998, 2001; Coste, 2001), and two "chronic skin disorders''-specific Ool instruments with French validated available versions, the VQ-Dermato (Grob et al., 1999, 2009) and the Skindex (Chren et al., 1996, 1997; Leplege et al., 2003). To better figure out the level of QoL in AA, we compared AA scores with those available in literature related to the French population: (1) rare dermatological diseases including hidradenitis suppurativa (Wolkenstein et al., 2007) and neurofibromatosis type 1 (Wolkenstein et al., 2001): (2) chronic/frequent dermatological diseases including psoriasis, chronic idiopathic urticaria, and atopic dermatitis (Grob et al., 2005); (3) general population: French age- and sex-matched controls (Leplege et al., 2001). This study was conducted in adherence to the Helsinki guidelines.

Institutional approval was not required for experiments. After having given their inform consent, 60 patients were included (39 women and 21 men); their mean age was 40.1 years (SD 15.2) and median AA duration was 6 years (2 months to 60 years). Course of the disease was stable in 25 subjects and unstable in 35. The median of the scalp surface involved was 77%. The median of severity score was 6.5 (range 4.0–9.0) by reference to the AA patients and 3.5 (range 2.0–6.0) by reference to the patients presenting any skin disorder.

Mental health and vitality were the most altered SF36 dimensions, whereas physical functioning, role physical and body pain were the least ones. Regarding VQ-Dermato, daily life, leisure activity, and physical discomfort were the least altered dimensions. For Skindex, emotions dimension was the most affected and symptoms the least one. Compared with the general population and with patients suffering from other dermatological conditions (Table 1), AA patients presented significantly altered Qol for almost all the SF36 dimensions. For VQ-Dermato, AA patients reported (i) significantly better (mood state, leisure activity, daily life, and physical discomfort) or worse scores (self-perception) than psoriasis, chronic idiopathic urticaria, and atopic dermatitis patients; (ii) being less bothered to treatment-induced restrictions than psoriasis, but more than chronic idio-

Abbreviations: AA, alopecia areata; Qol, quality of life; SF36, short form 36

	AA, <i>N</i> =60	HS, <i>N</i> =61	Р	NF1 <i>, N</i> =128	P	PSO, N=408	P	CU, N=367	<u>у</u> Р	AD, <i>N</i> =386	P	Controls	Р
SF36 <sup>2</sup>													
Physical functioning	88.2 ± 22.5	71.3 ± 27.4	< 0.001	$76.8 \pm 26.4$	0.004	L .						$90.3 \pm 7.0$	0.506
Role—physical	73.3 ± 35.0	43.6±39.0	< 0.001	72.8±39.1	0.932	2						87.0±6.7	0.006
Bodily pain	$77.2\pm20.7$	$44.5\pm24.4$	< 0.001	$65.3\pm29.6$	0.005	;						$77.9\pm6.3$	0.865
General health	$64.3 \pm 22.7$	$43.3 \pm 22.4$	< 0.001	$58.4 \pm 23.0$	0.101							$72.2 \pm 5.3$	0.011
Vitality	$54.5\pm20.4$	$40.4 \pm 20.4$	< 0.001	$49.7 \pm 21.3$	0.146							$62.4 \pm 3.2$	0.005
Social functioning	$58.9\pm29.5$	$52.5 \pm 25.8$	0.206	$70.4 \pm 25.7$	0.007	,						84.1 ± 3.5	< 0.001
Role—emotional	$64.1 \pm 39.1$	$42.9\pm42.0$	0.004	$69.4\pm39.4$	0.390	)						$86.3 \pm 5.1$	< 0.001
Mental health	49.3 ± 20.4	43.0±18.4	0.077	$56.4 \pm 22.0$	0.036	ò						69.7 ± 2.7	< 0.001
VQ-Dermato <sup>3</sup>													
Self-perception	51,4±26.2	$52.2 \pm 26.7$	0.868			37.4±24.7	< 0.001	23.8±21.8	< 0.001	$34.2 \pm 24.6$	< 0.001	I	
Daily life	$13.9 \pm 17.7$	$46.9\pm28.7$	< 0.001			$19.3 \pm 19.4$	0.145	$36.2 \pm 20.4$	< 0.001	$35.5 \pm 21.3$	< 0.001	I	
Mood state	$34.2 \pm 24.1$	$37.9 \pm 26.4$	0.422			49.3 ± 25.2	< 0.001	$50.3 \pm 25.5$	< 0.001	$50.1 \pm 25.5$	< 0.001	I	
Social functioning	$31.8 \pm 25.1$	$47.6\pm29.7$	0.002			31.3 ± 23.7	0.880	) 27.5 ± 22.9	0.184	34.1 ± 23.5	0.485	;	
Leisure activity	$23.9\pm28.4$	52.1 ± 31.3	< 0.001			$47.2 \pm 29.3$	< 0.001	36.7±28.1	< 0.001	$46.7 \pm 27.9$	< 0.001	I	
Treatment restrictions	$30.3 \pm 30.2$	$35.8\pm34.0$	0.349			$38.6 \pm 26.0$	0.024	17.0±20.7	< 0.001	$32.5 \pm 26.4$	0.556	5	
Physical discomfort	25.0±26.9	66.1 ± 26.5	< 0.001			44.4±28.2	< 0.001	61.4±23.7	< 0.001	69.8±21.3	< 0.001	I	
Skindex <sup>4</sup>													
Emotions	$48.9\pm27.8$	$59.2 \pm 23.4$	0.029	31.6±26.7	< 0.001								
Symptoms	$18.3 \pm 19.7$	$52.2 \pm 22.3$	< 0.001	$21.4 \pm 19.7$	0.315	5							
Functioning	$28.0 \pm 24.6$	48.8±25.5	< 0.001	$22.3 \pm 23.3$	0.126								

Table 1. Quality-of-life indicative comparisons between AA patients and other dermatological conditions, and French age- and sex-matched controls<sup>1</sup>

Abbreviations: AA, alopecia areata; AD, atopic dermatitis; CU, chronic urticaria; HS, hidradenitis suppurativa; NF1, neurofibromatosis type 1; PSO, psoriasis; SF36, short form 36.

<sup>1</sup>From Leplege *et al.*, (2001), Wolkenstein *et al.*, (2001), Grob *et al.*, (2005), and Wolkenstein *et al.*, (2007).

<sup>2</sup>SF36, 36 items, eight dimensions (range (0–100), 0 lowest and 100 highest level of Qol; Leplege *et al.*, 1998; Coste, 2001; Leplege *et al.*, 2001).

<sup>3</sup>VQ-Dermato, 28 items, seven domains and one overall score (range (0–100), 0 highest and 100 lowest level of Qol; Grob et al., 1999, 2009).

<sup>4</sup>Skindex, 29 items, three domains (range (0–100), 0 highest and 100 lowest level of Qol; Chren *et al.*, 1996, 1997; Leplege *et al.*, 2003). Bold values P < 0.05.

# Table 2. Associations between VQ-Dermato dimension scores and global score, and sociodemographic/clinical characteristics in 60 alopecia areata (AA) patients

	Self- perception	Daily life	Mood state	Social functioning	Leisure activity	Treatment restrictions	Physical discomfort	Global score
Gender <sup>1</sup>								
Men	$42.28 \pm 21.54$	$10.24 \pm 15.04$	$32.74 \pm 21.28$	$27.40 \pm 18.00$	$13.10 \pm 17.98$	$30.47 \pm 26.99$	$19.05 \pm 25.19$	$25.88 \pm 15.04$
Women	$53.66 \pm 28.47$	$16.05 \pm 18.94$	$35.02 \pm 25.78$	$34.29 \pm 28.23$	$30.07 \pm 31.51$	$30.15\pm32.00$	$28.38 \pm 27.58$	$32.30 \pm 21.83$
Р	0.196	0.341	0.839	0.499	0.042	0.694	0.161	0.406
Educational level <sup>1</sup>								
<12 Years	$48.58 \pm 28.48$	$13.75 \pm 18.57$	$27.08 \pm 23.99$	$26.09 \pm 25.18$	$21.21 \pm 29.06$	$26.14 \pm 29.61$	$24.46 \pm 27.30$	$27.02 \pm 21.57$
≥12 Years	$53.18 \pm 24.89$	$14.02 \pm 17.36$	$38.54 \pm 23.41$	$35.51 \pm 24.74$	$25.58 \pm 28.31$	$33.48\pm30.82$	$25.36 \pm 27.03$	$32.40 \pm 18.69$
Р	0.409	0.670	0.059	0.103	0.429	0.380	0.934	0.149

Table 2 continued on the following page

Table 2. Cont	inued							
	Self- perception	Daily life	Mood state	Social functioning	Leisure activity	Treatment restrictions	Physical discomfort	Global score
Marital status <sup>1</sup>								
Single	59.16±22.61	14.87±17.88	36.61 ± 25.72	38.65 ± 28.62	30.95 ± 32.07	35.16±38.52	26.25 ± 30.05	34.71 ± 24.22
Couple	47.09±27.33	13.43 ± 17.82	32.83 ± 23.36	28.07 ± 22.49	19.93 ± 25.74	$27.94 \pm 25.76$	24.34 ± 25.49	27.88 ± 17.33
Р	0.109	0.583	0.614	0.141	0.210	0.815	0.952	0.457
Occupational statu	's <sup>1</sup>							
Not working	43.63 ± 26.97	8.51 ± 12.41	28.08 ± 23.79	29.62 ± 25.49	23.02 ± 28.25	27.63 ± 31.62	21.59 ± 27.87	$27.43 \pm 21.63$
Working	$56.00 \pm 24.97$	17.06±19.63	37.67 ± 23.87	33.15±25.16	24.44 ± 28.90	31.85 ± 29.73	27.08 ± 26.47	31.66 ± 19.09
P	0.075	0.119	0.159	0.603	0.973	0.554	0.403	0.371
Disease course <sup>1,3</sup>								
Stable	$51.37 \pm 30.26$	$15.95 \pm 20.00$	$38.28 \pm 26.41$	$35.27 \pm 27.75$	$29.34 \pm 29.88$	$40.48 \pm 35.33$	$27.60 \pm 31.27$	34.88 ± 23.31
Unstable	$51.40 \pm 23.26$	$12.42 \pm 15.96$	31.31 ± 22.25	29.31 ± 23.11	$20.10 \pm 27.15$	$22.84 \pm 23.87$	$23.16 \pm 23.66$	27.07 ± 17.04
Р	0.701	0.630	0.340	0.474	0.138	0.093	0.792	0.174
Scalp surface invol	lved <sup>1</sup>							
<80%	$47.35 \pm 24.20$	$11.96 \pm 14.59$	$29.22 \pm 22.37$	$29.91 \pm 23.69$	$15.48 \pm 22.88$	$20.97 \pm 22.22$	$25.35 \pm 27.14$	25.34 ± 16.81
≥80%	$57.70 \pm 28.47$	17.01±21.76	$42.33 \pm 25.07$	$34.86 \pm 27.49$	$36.78 \pm 31.60$	$45.39 \pm 35.65$	24.43 ± 27.13	$38.20 \pm 22.43$
Р	0.065	0.693	0.064	0.512	0.006	0.016	0.887	0.023
Extrascalp involver	ment <sup>1</sup>							
No	$48.82 \pm 24.25$	$11.68 \pm 14.84$	30.34±21.39	$29.53 \pm 22.80$	$17.32 \pm 24.46$	$23.11 \pm 24.43$	$24.68 \pm 26.98$	$26.08 \pm 16.15$
Yes	$56.40 \pm 29.67$	$18.38 \pm 22.13$	$42.11 \pm 27.78$	$36.33 \pm 29.24$	$36.46 \pm 31.73$	$44.12 \pm 35.94$	$25.66 \pm 27.47$	$38.32 \pm 24.39$
Р	0.178	0.405	0.113	0.484	0.016	0.043	0.890	0.049
Alopecia universal	is <sup>1</sup>							
No	$50.55 \pm 25.30$	$14.86 \pm 17.46$	$32.82 \pm 22.97$	31.16±24.16	$21.85 \pm 27.48$	$29.38 \pm 30.16$	26.63 ± 27.21	30.30±19.16
Yes	$54.35 \pm 30.10$	$10.35 \pm 18.89$	$38.94 \pm 28.08$	$34.23 \pm 29.22$	$31.09 \pm 31.58$	$33.75 \pm 31.76$	$18.75 \pm 25.84$	$29.65 \pm 23.64$
Р	0.458	0.260	0.487	0.840	0.309	0.656	0.327	0.970
Age <sup>2</sup>	-0.099	-0.134	-0.236	-0.149	-0.173	-0.120	-0.059	-0.294
Р	0.456	0.322	0.075	0.260	0.195	0.406	0.661	0.042
Disease duration <sup>2</sup>	-0.118	-0.090	0.073	-0.157	0.084	-0.095	-0.145	-0.039
Р	0.376	0.511	0.589	0.239	0.536	0.518	0.281	0.793
Severity 1 <sup>2,4</sup>	0.247	0.027	0.319	0.180	0.359	0.396	-0.057	0.348
Р	0.059	0.844	0.015	0.173	0.006	0.004	0.673	0.015
Severity 2 <sup>2,5</sup>	0.216	0.056	0.334	0.187	0.383	0.377	0.038	0.390
Р	0.101	0.679	0.010	0.156	0.003	0.007	0.777	0.006

<sup>1</sup>Mean ± SD, *P*: *P*-value Mann–Whitney test.

<sup>2</sup>Spearman's correlation coefficient, *P. P*-value Spearman's test. <sup>3</sup>Course of the disease was defined as "unstable" if there was alternation of worsening and improvement phases in the last 2 years, and "stable" otherwise. <sup>4</sup>Severity 1: visual analog scale by reference to the cases of AA seen in daily practice; 0 "patient among the least affected" and 100 "patient among the most affected".

<sup>5</sup>Severity 2: visual analog scale by reference to the cases of all skin disorders seen in daily practice; 0 "patient among the least affected" and 100 "patient among the most affected".

Bold values: P < 0.05.

pathic urticaria patients. AA patients reported significantly better Qol than did hidradenitis suppurativa patients, except for social functioning and mental health dimensions of SF36 and self-perception, mood state, and treatment restriction dimensions of VQ-Dermato.

Skindex and SF36 were not statistically linked to sociodemographic and clinical parameters (data not shown). Only the VQ-Dermato global score indicated a significantly better Qol in older subjects (Table 2). Disease severity, extrascalp involvement, and scalp surface involved were related with altered VQ-dermato dimensions (Table 2).

Our results show that (1) Qol is impaired in AA, the most influenced domains being self-perception, such as in the Gulec et al (2004) study, and also mental health and social life. It may be because of the special importance of hair in appearance (Cash, 1999; Firooz et al., 2005); (2) social life is impaired in AA at the same level as in psoriasis, atopic dermatitis, and chronic idiopathic urticaria. On the dimensions dealing with mental health and social life, AA also compares very well with hidradenitis suppurativa, one of the rare skin disorders with the highest impact on most dimensions of Ool (Wolkenstein et al., 2007). Similarly, social life and mental comfort of patients seem to be more affected in AA than in neurofibromatosis type 1; (3) sociodemographic parameters did not impact Qol, except for leisure activities, with women appearing to be more affected; (4) clinical severity of AA appears to be poorly linked to Qol, even when assessed with a sensitive dermatology-specific Qol tool (VQ-Dermato). Furthermore, the dimension of self-perception, which seems to be particularly involved in AA, is not significantly linked to the clinical severity of the disorder.

Some limitations must be mentioned. The sample size did not allow a multivariate approach, and moderate associations were possibly missed owing to low power. We were unable to confirm the impact of anxiety or depression on Qol (Gulec *et al.*, 2004) because these parameters were not collected. One must be cautious while generalizing the study findings to minor cases seen in everyday care because our hospital-based series of AA was probably not representative of community cases. Although we did not systematically search for other health disorders in our patients, none of them declared to have one.

Although AA is a perfectly benign disorder, this work confirms the initial hypothesis that AA seriously impairs Qol, mainly by altering self-perception and self-esteem, both of which interfere with social life. From a practical point of view, these results can (i) help doctors to realize that they probably strongly underestimate the severity of AA and encourage them to give patients the treatments and the psychological help they require; (ii) help patients by showing that their suffering is understood by others; (iii) help health-care decision-makers to promote therapeutic trials in this orphan disorder.

#### **CONFLICT OF INTEREST**

The authors state no conflict of interest.

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#### REFERENCES

- Cash TF (1999) The psychosocial consequences of androgenetic alopecia: a review of the research literature. *Br J Dermatol* 141:398–405
- Chren MM, Lasek RJ, Flocke SA *et al.* (1997) Improved discriminative and evaluative capability of a refined version of Skindex, a quality-of-life instrument for patients with skin diseases. *Arch Dermatol* 133:1433–40
- Chren MM, Lasek RJ, Quinn LM *et al.* (1996) Skindex, a quality-of-life measure for patients with skin disease: reliability, validity, and responsiveness. *J Invest Dermatol* 107: 707–13
- Coste J (2001) Medical Outcome Study Short Form 36 Item Health Survey. Manual and Score Interpretation. Paris: Estem, 156pp
- Firooz A, Firoozabadi MR, Ghazisaidi B *et al.* (2005) Concepts of patients with alopecia areata about their disease. *BMC Dermatol* 5:1
- Grob JJ, Auquier P, Dreyfus I *et al.* (2009) How to prescribe antihistamines for chronic idiopathic urticaria: desloratadine daily versus PRN and quality of life. *Allergy* 64:605–12
- Grob JJ, Auquier P, Martin S *et al.* (1999) Development and validation of a quality of life measurement for chronic skin disorders in french: VQ-Dermato.The Reseau d' Epidemiolo gie en Dermatologie. *Dermatology* 199:213–22
- Grob JJ, Revuz J, Ortonne JP *et al.* (2005) Comparative study of the impact of chronic urticaria, psoriasis and atopic dermatitis on the quality of life. *Br J Dermatol* 152: 289–95
- Gulec AT, Tanriverdi N, Duru C *et al.* (2004) The role of psychological factors in alopecia areata and the impact of the disease on the quality of life. *Int J Dermatol* 43:352–6
- Leplege A, Ecosse E, Pouchot J et al. (2001) MOS SF36 Questionnaire. Manual and Guidelines for Scores' Interpretation. Vernouillet: Estem, 156
- Leplege A, Ecosse E, Verdier A *et al.* (1998) The French SF-36 Health Survey: translation, cultural adaptation and preliminary psychometric evaluation. *J Clin Epidemiol* 51:1013–23
- Leplege A, Ecosse E, Zeller J *et al.* (2003) The French version of Skindex (Skindex-France). Adaptation and assessment of psychometric properties. *Ann Dermatol Venereol* 130:177–83
- Wolkenstein P, Loundou A, Barrau K et al. (2007) Quality of life impairment in hidradenitis suppurativa: a study of 61 cases. J Am Acad Dermatol 56:621–3
- Wolkenstein P, Zeller J, Revuz J et al. (2001) Quality-of-life impairment in neurofibromatosis type 1: a cross-sectional study of 128 cases. Arch Dermatol 137:1421–5