Vertex balding, plasma insulin-like growth factor 1, and insulin-like growth factor binding protein 3

Elizabeth A. Platz, ScD,^a Michael N. Pollak, MD,^c Walter C. Willett, MD, DrPH,^{a,b,d} and Edward Giovannucci, MD, ScD^{a,b,d} Boston, Massachusetts, and Montreal, Canada

Background: A recent report suggested that men with vertex balding have higher levels of plasma insulin-

Methods: Participants were 431 male members of the Health Professionals Follow-up Study who responded to a question in 1992 on their hair pattern at 45 years of age and who were 47 to 81 years old when they provided a blood specimen in 1993 1994. Odds ratios (ORs) of vertex helding associated with

controlling for age at blood draw.

Results: Of the 431 men, 128 had vertex balding at age 45. Compared with men who were not balding, for a 1 standard deviation increase in plasma IGF-1 level (72.4 ng/mL), the OR for vertex balding was 1.31 (95% CI, 0.95-1.81). For a 1 standard deviation increase in plasma IGFBP-3 (957 ng/mL), the OR for vertex balding was 0.62 (95% CI, 0.44-0.88).

Conclusion: Older men with vertex balding have lower circulating levels of IGFBP-3 and higher levels of IGF-1 when controlling for IGFBP-3 level.

ale scalp hair pattern in adulthood has long been known to be influenced by andro-

study among 51 Greek men older than 65 years and found that men with vertex balding had higher cir-

tion of the hair follicle through its cycle of growth, senescence, and regeneration. Signorello et al² evaluated the relation of serum insulin-like growth factor 1 (IGF-1) and male pattern balding in a case-control

risk of vertex balding was 1.6 (95% confidence interval [CI], 0.9-3.2) after adjusting for age and 2.0 (95% CI, 1.0-4.6) after also adjusting for steroid hormone concentrations. Serum insulin-like growth factor

tem for 101-1, was not measured in the Oreck study.

Unit, Departments of Medicine and Uncology, Jewish General

Hospital and Harvard Medical School, Boston.d

Supported by research grants CA 55075 and HL 35464 from the
National Institutes of Health, Department of Health and Human
Services

reduced risk of vertex balding among middle-aged and elderly men.

MATERIAL AND METHODS

Participants for this analysis were selected from among members of the Health Professionals Follow-

Table 1. Plasma concentrations of IGF-1 and IGFBP-3 according to vertex balding* at age 45: Health Professionals Follow-up Study 1994

Diagna constituent	Vostor halding	Not vostov kaldina	A reluci-
No. of men	128	303	
Age at blood draw in 1994 (y)	64.7 ± 8.3	63.8 ± 8.2	.3
Unadjusted			
IGF-1 (ng/mL)	185.0 ± 71.4	190.2 ± 72.4	.5
IGFBP-3 (ng/mL)	3049 ± 871	3285 ± 957	.02
Mutually adjusted‡			
IGF-1 (ng/mL)	194.4 ± 44.4	186.2 ± 50.3	.11
IGFBP-3 (ng/mL)	3084 ± 536	3270 ± 664	.002

Data are presented as mean \pm standard deviation.

Modert, moderate or substantial vertex halding as self-reported using pictograms in 1992

Table II. Relation of vertex balding* with plasma IGF-1 and IGFBP-3: Health Professionals Follow-up Study 1994

		Tertile†				
	1	2	3	Unit‡	OR§	P value§
IGF-1						
Cases/controls	47/102	35/102	46/99			
Median (ng/mL)	123.1	181.9	251.2	72.4		
OR	1.00	1.00	1.86		1.31	.09
95% CI	Referent	0.56-1.78	0.93-3.70		0.95-1.81	
IGFBP-3						
Cases/controls	53/101	43/101	32/101			
Median (ng/mL)	2429	3195	4149	957		
OR	1.00	0.69	0.42		0.62	.008
OFW CL	Defenset	0 20 1 22	0.21.0.00		0.44-0.00	

^{*}Modest, moderate, or substantial vertex balding as self-reported using pictograms in 1992 versus no or little hair loss or receding hairline only.

IGF-1, we mutually adjusted for these two plasma levels and controlled for age at blood draw. Compared with men in the bottom tertile of IGF-1, the OR for vertex balding in the top tertile of IGF-1 was 1.86 (95% CI, 0.93-3.70) (Table II). Men in the top tertile of IGFBP-3 had a statistically significantly 58% lower risk of vertex balding than men in the bot-

receding hairline only (IGF-1: OR = 1.26, 95% CI [0.90-1.77]; IGFBP-3: OR = 0.61, 95% CI [0.42-0.89]). There was no evidence that the risk of balding associated with IGF-1 or IGFBP-3 was stronger with increasing extent of vertex balding at age 45 in an analysis limited only to men with vertex balding.

Because smoking may influence the IGF axis, we

IGF-1 of 72.4 ng/mL, the OR for vertex balding was 1.31 (95% CI, 0.95-1.81; P = .09). For a 957 ng/mL increase in IGFBP-3, the OR for vertex balding was 0.62 (95% CI, 0.44-0.88; P = .008) (Table II). These

increments). The OR for vertex balding modestly increased to 1.02 (95% CI, 0.57-1.84) and 2.00 (95% CI, 0.99-4.04) for the middle and top tertiles of IGF-1 and decreased to 0.66 (95% CI, 0.37-1.18) and 0.40

tFor comparison using the t test.

[‡]By residuals analysis.

[†]ORs from a logistic regression model with plasma level entered as two indicator variables and adjusted for age at blood draw (continuous). Tertile cut points for plasma levels of each factor were determined from the distribution of levels among the controls. ‡One standard deviation.

iiiviutuany aujusteu.

IGF-1 adjusted for IGFBP-3 was not as great as shown for IGF-1 by Signorello et al.² Differences in the two studies that might contribute to the disparity in the strength of the association between IGF-1 and vertex balding include different IGF-1 assays, older average age in the Greek study, interviewer-assessed balding in the Greek study versus self-report in our study, and IGF-1 and balding assessed concurrently in the Greek study versus 2 to 36 years apart in our study.

samples that we included in this analysis, adjustment did not appear to alter our estimates for the relation

these methodologic and population differences

We thank Alan Paciorek and Mira Kaufman at Harvard and Noreen Majeed and Yuzhen 'lao at the Jewish General Hospital for their laboratory and programming support.

REFERENCES

- 1. Demark-Wahnefried W, Lesko SM, Conaway MR, Robertson CN,
- 2. Signorello LB, Wuu J, Hsieh C-c, Tzonou A, Trichopoulos D, Mantzoros CS. Hormones and hair patterning in men: a role for insulin-like growth factor 1? J Am Acad Dermatol 1999;40:200-
- 3. Norwood OT. Male pattern baldness: classification and incidence. South Med J 1975;68:1359-65.
- 4. Olsen EA, Weiner MS, Delong ER, Pinnell SR. Topical minoxidil in

- 5. Rosner B. Fundamentals of biostatistics. Boston: Duxbury Press;

- - irriacology, pasic ariu cillical aspects, poca natoli, che

like growth factors on cultured human hair follicles: IGF-I at physiologic concentrations is an important regulator of hair follicie growth *in vitro.* J invest Dermatol 1994;102:857-61.

- 10. Peus D, Pittelkow MR. Growth factors in hair organ development and the hair growth cycle. Dermatol Clin 1996;4:559-72.
- 11. Itami S, Kurata S, Takayasu S. Androgen induction of follicular epithelial cell growth is mediated via insulin-like growth factors-1 from dermal papilla cells. Biochem Biophys Res Commun
- 12. Horton R, Pasupuletti V, Antonipillai I. Androgen induction of steroid 5 alpha-reductase may be mediated via insulin-like growth factor-I. Endocrinol 1993;133:447-51.
- 13. Kaufman KD, Olsen EA, Whiting D, Savin R, DeVillez R, Bergfeld W. et al. Finasteride in the treatment of men with androgenetic
- - Becker K, editor. Principles and practice of endocrinology and
- 16. Chan JM, Stampfer MJ, Giovannucci E, Gann PH, Ma J, Wilkinson

- 17. Mantzoros CS, Tzonou A, Signorello LB, Stampfer M,
- work A, Mantzoros CS, Andersson SW, Bergstrom R, Signorello LB, Lagiou P, et al. Insulin-like growth factor I and prostate cancer risk: a population-based, case-control study. J Nat! Cancer Inst 1998;90:911-5.
- 19. Peyrat JP, Bonneterre J, Hecquet B, Vennin P, Louchez MM, Fournier C, et al. Plasma insulin-like growth factor-I (IGF-I) concentrations in human breast cancer. Eur J Cancer 1993;29A:492-
 - 3 is decreased in early-stage operable pre-menopausal breast cancer. Int J Cancer 1995;62:266-70.
- 21. Hankinson SE, Willett WC, Colditz GA, Hunter DJ, Michaud DS, Deroo B, et al. Circulating concentrations of insulin-like growth factor-I and risk of breast cancer. Lancet 1998;351:1393-6.
- 22. Bohlke K, Cramer DW, Trichopoulos D, Mantzoros CS. Insulin-like growth factor-l in relation to premenopausal ductal carcinoma in situ of the breast. Epidemiology 1998;9:570-3.

plasma levels of insulin-like growth factor (IGF)-I and IGF-bind-