Prevalence of hypercortisolism in Type 2 Diabetes patients: a systematic review and meta-analysis

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Background: Type 2 Diabetes and Cushing's syndrome share clinical characteristics such as insulin resistance, hyperglycemia, hypertension, dyslipidemia and obesity. Several small studies have recorded a high prevalence of hypercortisolism in T2D. This could have therapeutic implications.

Objectives

 To perform a systematic review and meta-analysis on the prevalence and clinical characteristics of endogenous hypercortisolism in T2D patients.

Methods

- Original articles assessing the prevalence of endogenous hypercortisolism in T2D were eligible.
- A search was performed in SCOPUS, MEDLINE and EMBASE.
- Data were pooled in a random effects logistic regression model and reported with 95% confidence intervals (95% CI).

Hypercortisolism in diabetes type 2 patients Proportion with hypercortisolism (95% CI) 7.8 (3.9, 14.8) 12.1 (8.8, 16.4) 2.8 (1.1, 7.3) Hau Liu Hiroi 0.3 (0.0, 2.3) 2.7 (1.0, 7.0) 10.0 (5.3, 18.1) 0.6 (0.1, 4.0) 1.0 (0.1, 6.8) 7.8 (3.5, 16.3) 8.5 (5.4, 13.3) 3.5 (1.7, 7.0) Mullan 0.7 (0.3, 1.6) Terzolo 0.0 (0.0, 3.6) 3.3 (1.1, 9.8) Leibowitz 3.6 (3.0, 4.4) **TOTAL**

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	J	1	Number of	M/F	Number of patients with hypercortisolism	Test used, number of positives (%)						Pituitary	Adrenal	Estonio
		Author	diabetic participants			MSC	МРС	OD	LDDST	ACTH elevated	UFC	adenoma	adenoma	Ectopic adenoma
	1	Leibowitz <i>et al,</i> 1996	90 (100%)	26/64	3 (3,3%)	_	_	4 (4.4%)	3 (3,3%)	0 (0%)	3 (3,3%)	2 (2,2%)	1 (1,11%)	0 (0%)
	2	Catargi <i>et al,</i> 2003	200 (100%)	49/151	17 (8.5%)	2 (1%)	17 (8.5 %)	52 (26%)	_	_	17 (8.5%)	3 (1.5%)	8 (4%)	0 (0%)
	3	Chiodini <i>et al,</i> 2005	289 (100%)	124/165	35 (12.1%)	-	_	51 (17.6%)	_	5 (1.7%)	8 (2.7%)	4 (1.4%)	21 (7.1%)	2 (0.7%)
	4	Hau Liu <i>et a</i> l, 2005	141 (100%)	141/0	4 (2.8%)	28 (19.9%)	_	_	4 (2.8%)	3 2.1%)	1 (0.7%)	0 (0%)	0 (0%)	0 (0%)
	5	Caetano <i>et a</i> l, 2007	103 (100%)	34/69	8 (7.8%)	12 (11.7%)	_	29 (28.2%)	1 (0.9%)	2 (1.9%)	0 (0%)	1 (0.9%)	4 (3.9%)	0 (0%)
	6	Reimondo <i>et al,</i> 2007	100 (100%)	67/37	1 (1%)	_	1 (1%)	5 (5%)	1 (1%)	1 (1%)	1 (1%)	1 (1 %)	0 (0%)	0 (0%)
	7	Newsome <i>et a</i> l, 2008	171 (100%)	102/69	1 (0.6%)	_	_	31 (18.1%)	_	_	3 (1.8%)	0 (0%)	0 (0%)	0 (0%)
	8	Taniguchi <i>et al,</i> 2008	77 (100%)	47/30	6 (7.8%)	-	27 (35%)	7 (9%)	-	6 (7.8%)	-	2 (2.6%)	0 (0%)	0 (0%)
	9	Gagliardi et al, 2010	100 (100%)	60/40	0 (0%)	3 (3%)	0 (0%)	0 (0%)	_	0 (0%)	0 (0%)	_	_	_
1	10	Hiroi et al, 2010	304 (100%)	181/123	1 (0.3%)	_	_	1 (0.3%)	1 (0.3%)	1 (0.3%)	0 (%)	0 (0%)	14 (4.6%)	0 (0%)
1	11	Mullan et al, 2010	201 (100%)	119/91	7 (3.5%)	47 (23%)	_	7 (3.5%)	0 (0%)	_	0 (0%)	0 (0%)	0 (0%)	0 (0%)
1	12	Murakami et <i>al,</i> 2010	90 (100%)	53/37	9 (10%)	_	57 (63.3%)	9 ((10%)	_	0 (0%)	_	_	_	_
1	13	Mert <i>et al,</i> 2012	148 (100%)	35/113	4 (2.6%)	_	_	9 (6.2%)	4 (2.6%)	_	_	2 (1.4%)	2 (1.4%)	0 (0%)
1	14	Terzolo <i>et al,</i> 2012	813 (100%)	428/385	6 (0.7%)	_	_	40 (4.9)	6 (0.7%)	1 (0.12%)	_	0 (0%)	3 (0.4%)	_

Characteristics of included articles

Results

- Fourteen articles were included, with a total of 2827 T2D patients.
- The pooled prevalence of hypercortisolism was 3.6% [95% CI: 3.0-4.4].
- The prevalence did not differ between studies of unselected patients and patients selected based on presence
 of metabolic features such as obesity or poor glycemic control (p=0.41 from meta-regression).
- Imaging in patients with hypercortisolism (n=102) revealed adrenal tumors and pituitary tumors in 14 % and 52 %, respectively.

Conclusion

- Hypercortisolism is a relatively frequent finding in T2D.
- A substantial proportion of the cases also presented with adrenal or pituitary neoplasia, which could provide the basis for surgical intervention.
- These findings should not be ignored and controlled trials should be considered.

EP - 70

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