Preoperative octreotide treatment of acromegaly - long-term results of a randomized, controlled study

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BACKGROUND

Early postoperative results from the Preoperative Octreotide Treatment of Acromegaly (POTA) study has been published previously (1), and like later randomized studies we demonstrated a beneficial effect of presurgical treatment with somatostatin analogues (SSA) in GH-secreting macroadenomas when evaluated 3-4 months postoperatively.

However, concerns about a potential lingering effect of SSA and thereby potential false positive results have been raised.

OBJECTIVE

The objective of this study was to evaluate the effect of SSA pretreatment on the long-term surgical cure rates.

METHODS

Newly diagnosed patients were randomized to either 6 months preoperative treatment with octreotide LAR (n=32) or to direct surgery (n=30). Of them, 51 (26/25) had macroadenomas. The patients were evaluated one and five years postoperatively. Cure was defined as a normal IGF-1 level alone or combined with nadir GH < 2 mU/L during an oral glucose tolerance test.

All patients that had received additional acromegaly treatment were considered not cured.

RESULTS

The proportion of patients receiving postoperative acromegaly treatment was equal in the two groups both one (Table 1) and five years postoperatively.

	Preoperative SSA (n=31)	Direct surgery (n=30)
Medical therapy (N)	8	5
SSA	6	3
Pegvisomant	1	1
SSA + cabergoline	1	1
Radiation therapy (N)	1	4
Repeated surgery (N)	1	2
Total (N)	8	9

Table 1. Additional treatment for acromegaly until one year postoperative

1. Carlsen SM et al, JCEM 2008

RESULTS cont.

One year postoperative, when using the combined criteria for cure, 32 % of all adenomas and 38 % of macroadenomas were cured in the pretreatment group compared to 27 % and 24 % in the direct surgery group, respectively (Table 2). There was no significant difference between the treatment groups.

Treatment	According to IGF-1				According to IGF-1 and GH					
group	N	Cured	Not	Cure	p-	N	Cure	Not	Cure	p-
			cured	%	value		d	cured	%	value
All										
Pretreatment	31	12	19	39	0.87	31	10	21	32	0.63
Direct surgery	30	11	19	37	0.87	30	8	22	27	0.03
Microadenomas										
Pretreatment	5	0	5	0	0 17	5	0	5	0	0.44
Direct surgery	5	3	2	60	0.17	5	2	3	40	U. 44
Macroadenomas										
Pretreatment	26	12	14	46	0.30	26	10	16	38	0.27
Direct surgery	25	8	17	32	0.30	25	6	19	24	0.27

Table 2. Cure rates at 1 year postoperative

Five years following operation, the cure rate in the pretreatment group was 41 % in total and 40 % in macroadenomas, versus 31 % and 27 % in the direct surgery group, respectively (Table 3). There was no significant difference between the treatment groups.

Treatment	According to IGF-1				According to IGF-1 and GH					
group	N	Cured	Not	Cure	p-	N	Cured	Not	Cure	p-
			cured	%	value			cured	%	value
All										
Pretreatment	28	12	16	43	0.70	27	11	16	41	0.45
Direct surgery	28	11	17	39	0.79	26	8	18	31	0.45
Microadenomas										
Pretreatment	5	2	3	40	1.0	5	2	3	40	1.0
Direct surgery	5	3	2	60	1.0	4	2	2	50	1.0
Macroadenomas										
Pretreatment	23	10	13	43	0.55	22	9	13	41	0.24
Direct surgery	23	8	15	34	0.55	22	6	16	27	0.34

Table 3. Cure rates at 5 years postoperative

CONCLUSION

These long-term data from the POTA study does not prove a beneficial effect of SSA presurgical treatment.

However, in absolute numbers and in accordance with 3 months postoperative data from the POTA study, an approximately 50 % increase in cure rate is found for macroadenomas. Accordingly, we can not exclude that this is due to lack of power.

