## Appendix 2 (as supplied by authors): DBS Studies

Study	Study Type	Number/Type of Patients	Control group	Outcome
Subcallosal Cingulate				
Gyrus				
Holtzheimer et al.	Prospective	17 (10 MDD, 7 with bipolar	Single-blind, sham lead-in	At one year follow-up, remission and
2012 (1)	open label trial	II)	phase in all patients for 4	response rate of 36%. At 2 years,
	with sham		weeks after initial	remission rate of 58% and response
	lead-in phase		implantation. Second sham	rate of 92%. Remission and response
			phase following 24 weeks of	rates based on Hamilton Depression
			active stimulation; abandoned	Rating Scale (HDRS). Efficacy similar
			after 3 subjects due to	for MDD and bipolar patients
			significant deterioration in depressive symptoms after	
			stimulation was discontinued	
Lozano et al. 2012 (2)	Prospective	21 (MDD)	No control group	At 6 months follow-up, response rate
	open label trial	( /	учет в постава в	of 48%; at one year follow-up,
				response rate of 29%. Response
				measured by HDRS
Mayberg et al. 2005	Case series	5 (MDD, one patient with	No control group	Follow-up 6 months. 4/6 responders,
(3)		bipolar II)		2/6 remission as measured by HDRS
Kennedy et al. 2011 (4)	Case series	20 (MDD, one patients with	No control group	At last follow-up (3-6 years following
		bipolar II)		implantation, mean=3.5), response
				rate=64.3% and remission rate=
				42.9% (by HDRS). Considerable
				improvement in social functioning: 65% of patients engaged in work-
				related activity at last follow-up
				compared to 10% prior to DBS
Ventral				compared to 10% prior to 200
caudate/Ventral				
Striatum				
Malone et al. 2009 (5)	Case series	15 (MDD)	No control group	Follow-up from 6-51 months. 8/15
, ,				responders and 6/15 in remission at

Appendix to: Lipsman N, Sankar T, Downar J, et al. Neuromodulation for treatment-refractory major depressive disorder. *CMAJ* 2013. DOI:10.1503/cmaj.121317. Copyright © 2014 The Author(s) or their employer(s).

				last follow-up measured by Montgomery-Asberg Depression Scale (MADRS)
Nucleus Accumbens				
Schlaepfer et al. 2008 (6)	Case series	3 (MDD)	No control group	Double-blind changes to stimulation parameters and assessment. HDRS scores decreased with stimulation and increased with stimulation off.
Bewernick et al. 2010 (7)	Case series	10 (MDD)	No control group	At 12 months, 5/10 had achieved >50% reduction in HDRS scores (i.e., responders). Antidepressant, antianhedonic, and antianxiety effects observed
Inferior Thalamic				
Peduncle				
Jimenez et al. 2005 (8)	Case report	1 (MDD with comorbid bulimia nervosa and borderline personality disorder)	No control group	Double-blind assessment protocol following initial period of 8 months with "on" stimulation. No relapse of depressive symptoms with DBS turned off for 12 months. Sustained remission at 24 months with DBS on.

## References

- 1. Holtzheimer PE, Kelley ME, Gross RE, et al. Subcallosal cingu- late deep brain stimulation for treatment-resistant unipolar and bipolar depression. *Arch Gen Psychiatry* 2012;69:150-8.
- 2. Lozano AM, Giacobbe P, Hamani C, et al. A multicenter pilot study of subcallosal cingulate area deep brain stimulation for treatment-resistant depression. *J Neurosurg* 2012;116:315-22.
- 3. Mayberg HS, Lozano AM, Voon V, et al. Deep brain stimulation for treatment-resistant depression. *Neuron* 2005;45:651-60.

- 4. Kennedy SH, Giacobbe P, Rizvi SJ, et al. Deep brain stimulation ②for treatment-resistant depression: follow-up after 3 to 6 years. *Am J Psychiatry* 2011;168:502-10.
- 5. Malone DA Jr, Dougherty DD, Rezai AR, et al. Deep brain stim- ulation of the ventral capsule/ventral striatum for treatment- resistant depression. *Biol Psychiatry* 2009;65:267-75.
- 6. Schlaepfer TE, Cohen MX, Frick C, et al. Deep brain stimula- tion to reward circuitry alleviates anhedonia in refractory major depression. *Neuropsychopharmacology* 2008;33:368-77.
- 7. Bewernick BH, Hurlemann R, Matusch A, et al. Nucleus accum- bens deep brain stimulation decreases ratings of depression and anxiety in treatment-resistant depression. *Biol Psychiatry* 2010;67:110-6.
- 8. Jiménez F, Velasco F, Salin-Pascual R, et al. A patient with a resistant major depression disorder treated with deep brain stim- ulation in the inferior thalamic peduncle. *Neurosurgery* 2005;57:585-93.