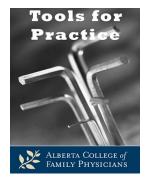
Tools for Practice is proudly sponsored by the Alberta College of Family Physicians (ACFP). ACFP is a provincial, professional voluntary organization, representing more than 4,000 family physicians, family medicine residents and medical students in Alberta. Established over fifty years ago, the ACFP strives for excellence in family practice through advocacy, continuing medical education and primary care research. www.acfp.ca

Reviewed: March 4, 2015 Evidence Updated: Added RCT and updated context Bottom line: No change

First Published: March 8, 2011



Vascular intervention for multiple sclerosis

Clinical Question: In patients with multiple sclerosis (MS), is angioplasty of obstructed extra-cranial venous lesions safe and does it improve MS symptoms?

Bottom-line: The initial study seemed promising, but was critically flawed for assessing benefit, and subsequent studies are generally not supportive. Presently, endovascular angioplasty for MS should not be recommended. The fluctuating nature of relapsing-remitting MS necessitates a long-term, multi-center, blinded RCT to determine if endovascular angioplasty is beneficial.

Evidence:

- Original cohort study, 65 MS patients with chronic cerebrospinal venous insufficiency (CCSVI) who all underwent angioplasty for obstructed azygous or internal jugular venous lesions.¹
 - Participants: Mean age 41, 46% male, minimal to moderate disability (not in wheelchair), taking MS disease-modifying agents.
 - Vascular outcomes: No serious operative or immediate post-operative complications.
 - Re-stenosis at 1-year ~50% for internal jugular.
 - Neurological outcomes at 18 months, compared to baseline (no control/placebo group):
 - No benefit seen in primary or secondary progressive MS subtypes.
 - Significant improvements in relapsing-remitting MS subtype.
 - Fewer patients relapsing over 1-year (50% versus 73%, p=0.0014).
 - Fewer patients with MRI lesions (12% versus 50%, p<0.0001).
 - Improved MS functional composite and quality of life scores.
 - Concerns: Single study site, not randomized, no control group, and unblinded.
 - Remission is a hallmark of relapsing-remitting MS: Untreated patients can have reductions (even prolonged) in clinical symptoms^{2,3} and/or MRI lesions.⁴
 A control group and long-term follow-up essential.

• Small double-blind RCT of internal jugular or azygous vein angioplasty found nonsignificantly higher relapse rates in the treatment arm (3/9 patients) versus the sham arm (1/10 patients) and higher rates of new MRI lesions in treatment group at six months.⁵

Context:

- Hypothesis of CCSVI and MS: Chronic insufficient cerebral venous drainage →
 cerebral iron deposits → engender the immune response underlying MS. Angioplasty
 improves cerebral blood outflow → decreases cerebral iron deposits → improves MS
 symptoms.⁶
- Many patients are having this unproven procedure, and serious complications are being observed.^{7,8}
- Subsequent studies question the relationship between CCSVI and MS.⁹⁻¹⁴
- Provider organizations and patient groups state there is no indication of a connection between CCSVI and MS and there are risks associated with angioplasty therapy. 15,16

Original Authors:

Michael R Kolber MD CCFP, Ken Makus MD FRCPC, G Michael Allan MD CCFP

Updated: Reviewed:

Braiden Hellec BSc Pharm G Michael Allan MD CCFP

References:

- 1. Zamboni P, Galeotti R, Menegatti E, et al. J Vasc Surg. 2009; 50:1348-58.
- 2. Martinez-Yelamos S, Martinez-Yelamos A, Martin-Ozaeta G, et al. Mult Scler. 2006; 12:826-9.
- 3. PRISMS Study Group. Lancet. 1998; 352:1498-504.
- 4. Zhao Y, Traboulsee A, Petkau AJ, et al. Neurology. 2008; 70:1092-7.
- 5. Siddiqui AH, Zivadinov R, Benedict RHB, et al. Neurology. 2014; 83:441-9.
- 6. Singh AV, Zamboni P. J Cereb Blood Flow Metab. 2009; 29:1867-78.
- 7. Samson K. Ann Neurol. 2010; 67(1):A13-5.
- 8. Kuehn BM. JAMA. 2012; 307:2575-6.
- 9. Doepp F, Paul F, Valdueza JM, et al. Ann Neurol. 2010; 68:173-83.
- 10. Sundstrom P, Wahlin A, Ambarki K, et al. Ann Neurol. 2010; 68:255-9.
- 11. Baracchini C, Perini P, Calabrese M, et al. Ann Neurol. 2011; 69:90-9.
- 12. Zivadinov R, Marr K, Cutter G, et al. Neurology. 2011; 77(2):138-44.
- 13. Tsivgoulis G, Sergentanis TN, Chan A, et al. Ther Adv Neurol Disord. 2014; 7(2):114-36.
- 14. Laupacis A, Lillie E, Dueck A, et al. CMAJ. 2011; 183(16):E1203-12.
- 15. Multiple Sclerosis Society of Canada. Chronic Cerebrospinal Venous Insufficiency (CCSVI). Available from: https://beta.mssociety.ca/research-news/hot-topics-in-research/chronic-cerebrospinal-venous-insufficiency-ccsvi. Accessed March 3, 2015.
- 16. National Multiple Sclerosis Society. CCSVI and MS. Available from: http://www.nationalmssociety.org/Research/Research-News-Progress/Rese

Tools for Practice is a biweekly article summarizing medical evidence with a focus on topical issues and practice modifying information. It is coordinated by G. Michael Allan, MD, CCFP and the content is written by practicing family physicians who are joined occasionally by a health professional from another medical specialty or health discipline. Each article is peer-reviewed, ensuring it maintains a high standard of quality, accuracy, and academic integrity.

The ACFP has supported the publishing and distribution of the Tools for Practice library since 2009. If you are not a member of the ACFP and would like to receive the TFP emails, please sign up for the distribution list at http://bit.ly/signupfortfp. Archived articles are available at no extra cost on the ACFP website.

You can now earn credits on Tools for Practice! In August 2014, the ACFP launched GoMainpro, an online accreditation tool to help facilitate MAINPRO® accreditation for the ACFP's Tools for Practice library which has been accredited for Mainpro-M1 credits by the College of Family Physicians of Canada (CFPC). The combination of the CFPC's Direct Entry Program and GoMainpro's tracking and reporting features provide an easy and convenient way to earn Mainpro-M1 credits.

This communication reflects the opinion of the authors and does not necessarily mirror the perspective and policy of the Alberta College of Family Physicians.