



TITLE: Anticholinergic Drugs for the Treatment of Overactive Bladder in Older Patients: Clinical Evidence, Safety, and Guidelines

DATE: 05 March 2012

RESEARCH QUESTIONS

1. What is the clinical evidence regarding any changes in cognitive function associated with the use of anticholinergic drugs for the treatment of overactive bladder in patients older than 65 years of age?
2. What is the clinical evidence regarding the safety of anticholinergic drugs, with or without cholinesterase inhibitors, for the treatment of overactive bladder in patients older than 65 years of age?
3. What are the evidence-based guidelines regarding the use of anticholinergic drugs for the treatment of overactive bladder in patients older than 65 years of age?

KEY MESSAGE

Seven clinical studies and one evidence-based guideline were identified regarding the use of anticholinergic drugs for the treatment of overactive bladder in patients older than 65 years of age.

METHODS

A limited literature search was conducted on key resources including PubMed, The Cochrane Library (2012, Issue 2), University of York Centre for Reviews and Dissemination (CRD) databases, Canadian and abbreviated list of major international health technology agencies, as well as a focused Internet search. No filters were applied to limit the retrieval by study type. Where possible, retrieval was limited to the human population. The search was also limited to English language documents published between January 1, 2007 and February 21, 2012. Internet links were provided, where available.

Disclaimer: The Rapid Response Service is an information service for those involved in planning and providing health care in Canada. Rapid responses are based on a limited literature search and are not comprehensive, systematic reviews. The intent is to provide a list of sources of the best evidence on the topic that CADTH could identify using all reasonable efforts within the time allowed. Rapid responses should be considered along with other types of information and health care considerations. The information included in this response is not intended to replace professional medical advice, nor should it be construed as a recommendation for or against the use of a particular health technology. Readers are also cautioned that a lack of good quality evidence does not necessarily mean a lack of effectiveness particularly in the case of new and emerging health technologies, for which little information can be found, but which may in future prove to be effective. While CADTH has taken care in the preparation of the report to ensure that its contents are accurate, complete and up to date, CADTH does not make any guarantee to that effect. CADTH is not liable for any loss or damages resulting from use of the information in the report.

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The summary of findings was prepared from the abstracts of the relevant information. Please note that data contained in abstracts may not always be an accurate reflection of the data contained within the full article.

RESULTS

Rapid Response reports are organized so that the higher quality evidence is presented first. Therefore, health technology assessment reports, systematic reviews, and meta-analyses are presented first. These are followed by randomized controlled trials, non-randomized studies, and evidence-based guidelines.

Two randomized controlled trials, five non-randomized studies, and one guideline were identified regarding the use of anticholinergic drugs for the treatment of overactive bladder in patients older than 65 years of age. No relevant health technology assessments, systematic reviews, or meta-analyses were identified. Additional references of potential interest are provided in the appendix.

OVERALL SUMMARY OF FINDINGS

No major changes to cognitive function, or safety issues, related to anticholinergic drugs were reported in any of the included studies.¹⁻⁷ Further details of the included studies are provided in Table 1.

Table 1: Summary of Included Studies

| Authors | Study Type and Patient Population | Interventions | Results |
|-------------------------------|--|------------------------------|---|
| Herschorn et al. ¹ | RCT Older adults with OAB | solifenacin or oxybutynin IR | The incidence of dry mouth and discontinuation of treatment was lower for patients receiving solifenacin. |
| Lackner et al. ² | RCT Women aged 65 years and older with urge incontinence and cognitive impairment | oxybutynin ER or placebo | In older women with dementia, there were no reports of delirium and no differences were found in confusion assessment scores, when compared with placebo. Oxybutynin was well tolerated. |
| Wawruch et al. ³ | NRS Hospitalized patients aged 65 years and older | anticholinergic medications | The authors identified urinary incontinence as a risk factor for the use of anticholinergic drugs and indicated that physicians should be mindful of potential adverse anticholinergic effects in elderly patients. |
| Gomes et al. ⁴ | NRS Adults aged 66 years and older being treated for OAB | oxybutynin IR or tolterodine | No difference in risk for falls was identified between the two groups. There was no increase in the risk of fracture or delirium associated with oxybutynin IR when compared with tolterodine. |
| Staskin et al. ⁵ | NRS Cognitively intact adults aged 65 to 75 | tropium ER | After 10 days of treatment, tropium was not detected in cerebrospinal fluid and no negative cognitive effects were observed. |

Table 1: Summary of Included Studies

| Authors | Study Type and Patient Population | Interventions | Results |
|--------------------------|--|--|---|
| | years with OAB | | |
| Isik et al. ⁶ | NRS Elderly patients with late onset Alzheimer's disease | tropium, galantamine, or tropium + galantamine | Patient satisfaction increased in both the tropium and combination therapy groups. Cognitive assessment scores did not change significantly during the six month course of treatment. The authors suggested that tropium, could safely be used for the treatment of overactive bladder in combination with cholinesterase inhibitors. |
| Sink et al. ⁷ | NRS Nursing home patients aged 65 years and older taking a cholinesterase inhibitor | oxybutynin IR + cholinesterase inhibitors or tolterodine + cholinesterase inhibitors | A 50 percent greater rate in quarterly active daily living functional decline was observed in higher-functioning patients using combination therapy compared with those treated with cholinesterase inhibitors alone. |

ER = extended-release; IR = immediate release;; NRS = non-randomized studies; OAB = overactive bladder; RCT = randomized controlled trial

The included evidence-based guideline⁸ indicates that the most common AEs experienced by patients using anticholinergic medications include dry mouth, blurred vision, dizzy spells, constipation, and urinary retention. Central nervous system effects such as cognitive disorders and confusion may also be experienced and may be of greatest concern for elderly patients. It is suggested that the efficacy of drugs for OAB be assessed at regular intervals.

REFERENCES SUMMARIZED

Health Technology Assessments

No literature identified.

Systematic Reviews and Meta-analyses

No literature identified.

Randomized Controlled Trials

1. Herschorn S, Pommerville P, Stothers L, Egerdie B, Gajewski J, Carlson K, et al. Tolerability of solifenacin and oxybutynin immediate release in older (> 65 years) and younger (<= 65 years) patients with overactive bladder: sub-analysis from a Canadian, randomized, double-blind study. *Curr Med Res Opin.* 2011 Feb;27(2):375-82.
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[PubMed: PM18410326](#)

Non-Randomized Studies

3. Wawruch M, Macugova A, Kostkova L, Luha J, Dukat A, Murin J, et al. The use of medications with anticholinergic properties and risk factors for their use in hospitalised elderly patients. *Pharmacoepidemiol Drug Saf.* 2012 Feb;21(2):170-6.
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4. Gomes T, Juurlink DN, Ho JM, Schneeweiss S, Mamdani MM. Risk of serious falls associated with oxybutynin and tolterodine: a population based study. *J Urol.* 2011 Oct;186(4):1340-4.
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[PubMed: PM18384584](#)

Guidelines and Recommendations

8. Finnish Medical Society Duodecim. Urinary incontinence in women. In: EBM Guidelines. Evidence-Based Medicine [Internet]. Helsinki (FIN): Wiley Interscience, John Wiley & Sons; 2008 Aug 8 [cited 2012 Feb 29].
National Guideline Clearinghouse summary available from:
<http://www.guideline.gov/content.aspx?id=13195>
See: Pharmacotherapy

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APPENDIX – FURTHER INFORMATION:

Systematic Reviews and Meta-Analyses

9. Paquette A, Gou P, Tannenbaum C. Systematic review and meta-analysis: do clinical trials testing antimuscarinic agents for overactive bladder adequately measure central nervous system adverse events? *J Am Geriatr Soc.* 2011 Jul;59(7):1332-9.
[PubMed: PM21718264](#)

Review Articles

10. Chancellor M, Boone T. Anticholinergics for overactive bladder therapy: central nervous system effects. *CNS Neurosci Ther.* 2012 Feb;18(2):167-74.
[PubMed: PM22070184](#)
11. Gerretsen P, Pollock BG. Drugs with anticholinergic properties: a current perspective on use and safety. *Expert Opin Drug Saf.* 2011 Sep;10(5):751-65.
[PubMed: PM21635190](#)
12. Oefelein MG. Safety and tolerability profiles of anticholinergic agents used for the treatment of overactive bladder. *Drug Saf.* 2011 Sep 1;34(9):733-54.
[PubMed: PM21830836](#)
13. Pagoria D, O'Connor RC, Guralnick ML. Antimuscarinic drugs: review of the cognitive impact when used to treat overactive bladder in elderly patients. *Curr Urol Rep.* 2011 Oct;12(5):351-7.
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[PubMed: PM18660862](#)
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[PubMed: PM18643858](#)
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