

Multiple Sclerosis

Appendices

Clinical Guideline <...>

Appendix F - LSS

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Draft for Consultation

*Commissioned by the National Institute for
Health and Care Excellence*

Disclaimer

Healthcare professionals are expected to take NICE clinical guidelines fully into account when exercising their clinical judgement. However, the guidance does not override the responsibility of healthcare professionals to make decisions appropriate to the circumstances of each patient, in consultation with the patient and/or their guardian or carer.

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Contents

Appendices.....	5
Appendix A: Literature search strategies	5

Appendices

Appendix A: Literature search strategies

Contents

Introduction	Search methodology
Section A.1	Population search strategies
A.1.1	Multiple sclerosis
A.1.2	Multiple sclerosis with clinically isolated syndrome
Section A.2	Study filter terms
A.2.1	Systematic reviews (SR)
A.2.2	Randomized controlled trials (RCT)
A.2.3	Observational studies (OBS)
A.2.4	Diagnostic accuracy (DA)
A.2.5	Patient information (PI)
A.2.6	Health economic studies (HE)
A.2.7	Quality of life studies (QOL)
Section A.3	Searches for specific questions with intervention
A.3.1	Diagnostic criteria
A.3.2	Information and support
A.3.3	Modifiable risk factors
A.3.4	Structured assessment
A.3.5	Coordination of care
A.3.6	Management of acute relapses
A.3.7	Pharmacological treatment of spasticity
A.3.8	Pharmacological treatment of mobility
A.3.9	Pharmacological treatment of fatigue
A.3.10	Pharmacological treatment of oscillopsia
A.3.11	Pharmacological treatment of emotionalism
A.3.12	Pharmacological treatment of ataxia and tremor
A.3.13	Non pharmacological treatment of memory and cognition
A.3.14	Non pharmacological treatment of ataxia and tremor
A.3.15	Non pharmacological treatment of fatigue
	Non pharmacological treatment of mobility
	Non pharmacological treatment of pain
	Non pharmacological treatment of spasticity
	Non pharmacological treatment of rehabilitation settings

Introduction	Search methodology
A.3.16	Vitamin D
A.3.17	Omegas
	Acupuncture
Section A.4	Health economic searches
A.4.1	Economic reviews
A.4.2	Quality of life reviews
Section A.5	References

Search strategies used for the Multiple Sclerosis guideline are outlined below and were run in accordance with the methodology in the NICE Guidelines Manual 2009:

http://www.nice.org.uk/media/615/64/The_guidelines_manual_2009.pdf. All searches were run up to **27th January 2014** unless otherwise stated. Any studies added to the databases after this date were not included unless specifically stated in the text. Where possible searches were limited to retrieve material published in English.

Table 1: Database date parameters

Database	Dates searched
Medline	1946 – 27 th January 2014
Embase	1980 – February 2014 (week 6)
The Cochrane Library	Cochrane Reviews to 2014 Issue 1 of 12 CENTRAL to 2013 Issue 12 of 12 DARE, HTA and NHSEED to 2013 Issue 4 of 4

Searches for the **clinical reviews** were run in Medline (OVID), Embase (OVID) and the Cochrane Library (Wiley). Additional searches were run in PsycINFO & Allied and Complementary Medicine (AMED) for some questions. Usually, searches were constructed in the following way:

- A PICO format was used for **intervention** searches where population (P) terms were combined with Intervention (I) and sometimes Comparison (C) terms. An intervention can be a drug, a procedure or a diagnostic test. Outcomes (O) are rarely used in search strategies for interventions. Search Filters were also added to the search where appropriate.
- A PEO format was used for **prognosis** searches where population (P) terms were combined with exposure (E) terms and sometimes outcomes (O). Search filters were added to the search where appropriate.

Searches for **patient information** were run in Medline (Ovid), Embase (Ovid), Cinahl (EBSCO) and PsychINFO (Ovid). Searches were constructed by adding a patient information search filter to the population terms.

Searches for the **health economic reviews** were run in Medline (Ovid), Embase (Ovid), the NHS Economic Evaluations Database (NHS EED), the Health Technology Assessment (HTA) database and the Health Economic Evaluation Database (HEED). Searches in NHS EED and HEED were constructed only using population terms. For Medline and Embase an economic filter (instead of a study type filter) was added to the same clinical search strategy. An additional search was run for Balofen.

A.1 Population search strategies

A.1.1 Multiple sclerosis population

Medline search terms

1.	exp *Multiple Sclerosis/
2.	((multiple or disseminated) adj2 scleros*).ti,ab.
3.	encephalomyelitis disseminata.ti,ab.
4.	MS.ti.
5.	Myelitis, Transverse/
6.	transverse myelitis.ti,ab.
7.	or/1-6

Embase search terms

1.	exp *Multiple Sclerosis/
2.	((multiple or disseminated) adj2 scleros*).ti,ab.
3.	encephalomyelitis disseminata.ti,ab.
4.	MS.ti.
5.	myelitis/
6.	transverse myelitis.ti,ab.
7.	or/1-6

Cochrane search terms

#1.	MeSH descriptor: [Multiple Sclerosis] explode all trees
#2.	multiple scleros* or disseminat* scleros*:ti,ab,kw
#3.	encephalomyelitis disseminata:ti,ab,kw
#4.	MS:ti
#5.	MeSH descriptor: [Myelitis, Transverse] explode all trees
#6.	Transverse Myelitis:ti,ab,kw
#7.	#1 or #2 or #3 or #4 or #5 or #6

Psycinfo search terms

1.	exp Multiple Sclerosis/
2.	((multiple or disseminated) adj2 scleros*).ti,ab.
3.	encephalomyelitis disseminata.ti,ab.
4.	MS.ti.
5.	transverse myelitis.ti,ab.
6.	*Myelitis/
7.	*demyelination/
8.	(demyelinat\$ adj2 (syndrome or disease or disorder)).ti,ab.
9.	or/1-8

AMED search terms

1.	multiple sclerosis/
2.	((multiple or disseminated) adj2 scleros\$).ti,ab.
3.	encephalomyelitis disseminata.ti,ab.
4.	(MS not "LC-MS").ti.
5.	transverse myelitis.ti,ab.

6.	or/1-5
7.	Models biological/ or exp rodents/
8.	(rat or rats or mice or mouse).ti.
9.	7 or 8
10.	Case report/ or case reports.pt.
11.	(letter or comment\$).ti.
12.	Randomized controlled trials/ or random\$.ti,ab.
13.	10 or 11
14.	13 not 12
15.	9 or 14
16.	6 not 15

A.1.2 Multiple sclerosis including clinically isolated syndrome (CIS)

Medline search terms MS

1.	exp *Multiple Sclerosis/
2.	((multiple or disseminated) adj2 scleros*).ti,ab.
3.	encephalomyelitis disseminata.ti,ab.
4.	MS.ti.
5.	Myelitis, Transverse/
6.	transverse myelitis.ti,ab.
7.	or/1-6
8.	*Demyelinating Diseases/
9.	*Demyelinating Autoimmune
10.	(Demyelinat* adj2 (syndrome or disease)).ti,ab.
11.	Chronic Cerebrospinal Venous Insufficiency.ti,ab.
12.	Venous Insufficiency/cf, co, di, et, us [Cerebrospinal Fluid, Complications, Diagnosis, Etiology, Ultrasonography]
13.	(Devic* adj (disease or syndrome)).ti,ab.
14.	(clinically isolated adj syndrome).ti,ab.
15.	exp Optic Neuritis/
16.	Neuromyelitis Optica.sh.
17.	Or/1-16

Embase search terms

1.	exp *Multiple Sclerosis/
2.	((multiple or disseminated) adj2 scleros*).ti,ab.
3.	encephalomyelitis disseminata.ti,ab.
4.	MS.ti.
5.	myelitis/
6.	transverse myelitis.ti,ab.
7.	*Demyelinating Diseases/
8.	*Demyelinating Autoimmune Diseases, CNS/
9.	(Demyelinat* adj2 (syndrome or disease)).ti,ab.
10.	Chronic Cerebrospinal Venous Insufficiency.ti,ab.
11.	Venous Insufficiency/cf, co, di, et, us [Cerebrospinal Fluid, Complications, Diagnosis, Etiology, Ultrasonography]

12.	(Devic* adj (disease or syndrome)).ti,ab.
13.	(clinically isolated adj syndrome).ti,ab.
14.	exp Optic Neuritis/
15.	Or/1-15

Cochrane search terms

#1.	MeSH descriptor: [Multiple Sclerosis] explode all trees
#2.	multiple scleros* or disseminat* scleros*:ti,ab,kw
#3.	encephalomyelitis disseminata:ti,ab,kw
#4.	MS:ti
#5.	MeSH descriptor: [Myelitis, Transverse] explode all trees
#6.	Transverse Myelitis:ti,ab,kw
#7.	MeSH descriptor: [Demyelinating Diseases] this term only
#8.	MeSH descriptor: [Demyelinating Autoimmune Diseases, CNS] this term only
#9.	(Demyelinati* near/2 (syndrome* or disease* or nerve*)):ti,ab,kw
#10.	Chronic Cerebrospinal Venous Insufficiency:ti,ab,kw
#11.	MeSH descriptor: [Venous Insufficiency] explode all trees and with qualifier(s): [Cerebrospinal fluid - CF, Complications - CO, Diagnosis - DI, Etiology - ET, Ultrasonography - US]
#12.	(Devic* near/2 (disease* or syndrome*)):ti,ab,kw
#13.	("clinically isolated syndrome" or "clinically isolated syndromes"):ti,ab,kw
#14.	MeSH descriptor: [Optic Neuritis] explode all trees
#15.	Neuromyelitis Optica:ti,ab,kw
#16.	{or #1-#15}

Psycinfo search terms

1.	exp Multiple Sclerosis/
2.	((multiple or disseminated) adj2 scleros*).ti,ab.
3.	encephalomyelitis disseminata.ti,ab.
4.	MS.ti.
5.	transverse myelitis.ti,ab.
6.	*Myelitis/
7.	*demyelination/
8.	(demyelinat\$ adj2 (syndrome or disease or disorder)).ti,ab.
9.	chronic cerebrospinal venous insufficiency.ti,ab.
10.	(devic\$ adj (disease or syndrome)).ti,ab.
11.	(clinically isolated adj syndrome).ti,ab.
12.	(neuromyelitis optica or optic neuritis).ti,ab.
13.	or/1-12

A.2 Study filter search terms**A.2.1 Systematic review (SR) search terms****Medline search terms**

1.	meta-analysis/
2.	meta-analysis as topic/
3.	(meta analy* or metanaly* or metaanaly* or meta regression).ti,ab.

4.	((systematic* or evidence*) adj2 (review* or overview*)).ti,ab.
5.	(reference list* or bibliograph* or hand search* or manual search* or relevant journals).ab.
6.	(search strategy or search criteria or systematic search or study selection or data extraction).ab.
7.	(search* adj4 literature).ab.
8.	(medline or pubmed or cochrane or embase or psychlit or psyclit or psychinfo or psycinfo or cinahl or science citation index or bids or cancerlit).ab.
9.	cochrane.jw.
10.	((multiple treatment* or indirect or mixed) adj2 comparison*).ti,ab.
11.	or/1-10

Embase search terms

1.	systematic review/
2.	meta-analysis/
3.	(meta analy* or metanaly* or metaanaly* or meta regression).ti,ab.
4.	((systematic or evidence) adj2 (review* or overview*)).ti,ab.
5.	(reference list* or bibliograph* or hand search* or manual search* or relevant journals).ab.
6.	(search strategy or search criteria or systematic search or study selection or data extraction).ab.
7.	(search* adj4 literature).ab.
8.	(medline or pubmed or cochrane or embase or psychlit or psyclit or psychinfo or psycinfo or cinahl or science citation index or bids or cancerlit).ab.
9.	cochrane.jw.
10.	((multiple treatment* or indirect or mixed) adj2 comparison*).ti,ab.
11.	or/1-10

Psycinfo search terms

1.	meta-analysis/
2.	(meta-analys\$ or metanalys\$ or metaanalys\$ or meta analys\$).ti,ab.
3.	((systematic or evidence\$ or methodol\$ or quantitativ\$) adj3 (review\$ or overview\$)).ti,ab.
4.	((pool\$ or combined or combining) adj2 (data or trials or studies or results)).ti,ab.
5.	(systematic or meta\$).pt. or (meta analysis or systematic review).md.
6.	or/1-6

A.2.2 Randomised controlled studies (RCTs) search terms**Medline search terms**

1.	randomized controlled trial.pt.
2.	controlled clinical trial.pt.
3.	randomi#ed.ab.
4.	placebo.ab.
5.	randomly.ab.
6.	clinical trials as topic.sh.
7.	trial.ti.
8.	or/1-7

Embase search terms

1.	random*.ti,ab.
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2.	factorial*.ti,ab.
3.	(crossover* or cross over*).ti,ab.
4.	((doubl* or singl*) adj blind*).ti,ab.
5.	(assign* or allocat* or volunteer* or placebo*).ti,ab.
6.	crossover procedure/
7.	double blind procedure/
8.	single blind procedure/
9.	randomized controlled trial/
10.	or/1-9

Psycinfo search terms

1.	exp clinical trial/
2.	randomi\$.ti,ab.
3.	((clinical or control\$) adj3 trial\$).ti,ab.
4.	((singl\$ or doubl\$ or trebl\$ or tripl\$) adj5 (blind\$ or mask\$)).ti,ab.
5.	(volunteer\$ or control group or controls).ti,ab.
6.	placebo/ or placebo\$.ti,ab.
7.	or/58-63

A.2.3 Observational studies (OBS) search terms**Medline search terms**

1.	epidemiologic studies/
2.	exp case control studies/
3.	exp cohort studies/
4.	cross-sectional studies/
5.	case control.ti,ab.
6.	(cohort adj (study or studies or analys*)).ti,ab.
7.	((follow up or observational or uncontrolled or non randomi#ed or nonrandomi#ed or epidemiologic*) adj (study or studies)).ti,ab.
8.	((longitudinal or retrospective or prospective or cross sectional) and (study or studies or review or analys* or cohort*)).ti,ab.
9.	or/1-8

Embase search terms

1.	clinical study/
2.	exp case control study/
3.	family study/
4.	longitudinal study/
5.	retrospective study/
6.	prospective study/
7.	cross-sectional study/
8.	cohort analysis/
9.	follow-up/
10.	cohort*.ti,ab.
11.	9 and 10
12.	case control.ti,ab.

13.	(cohort adj (study or studies or analys*)).ti,ab.
14.	((follow up or observational or uncontrolled or non randomi#ed or nonrandomi#ed or epidemiologic*) adj (study or studies)).ti,ab.
15.	((longitudinal or retrospective or prospective or cross sectional) and (study or studies or review or analys* or cohort*)).ti,ab.
16.	or/1-8,11-15

Cochrane search terms

#1.	MeSH descriptor: [Cohort Studies] explode all trees
#2.	MeSH descriptor: [Cross-Sectional Studies] explode all trees
#3.	((prospective or "cross sectional" or "follow up" or longitudinal or comparative or cohort*) and (study or studies or review or analys*)):ti,ab
#4.	"Comparative study":pt
#5.	#1 or #2 or #3 or #4

A.2.4 Patient Information (PI) search terms**Medline search terms**

1.	exp empirical research/
2.	QUESTIONNAIRES/
3.	(questionnaire\$ or survey\$).ti,ab.
4.	Health Surveys/
5.	Health Care Surveys/
6.	Interview/
7.	exp Interviews as Topic/
8.	interview\$.tw.
9.	grounded theory.tw.
10.	exp Nursing Methodology Research/
11.	focus group\$.tw.
12.	themes.tw.
13.	phenomenology.tw.
14.	ethnograph\$.tw.
15.	qualitative research/
16.	qualitative.tw.
17.	or/1-16

Embase search terms

1.	exp empirical research/
2.	QUESTIONNAIRES/
3.	(questionnaire\$ or survey\$).ti,ab.
4.	Health Surveys/
5.	Health Care Surveys/
6.	Interview/
7.	exp Interviews as Topic/
8.	interview\$.tw.
9.	grounded theory.tw.
10.	exp Nursing Methodology Research/

11.	focus group\$.tw.
12.	themes.tw.
13.	phenomenology.tw.
14.	ethnograph\$.tw.
15.	qualitative research/
16.	qualitative.tw.
17.	or/1-16

PsychInfo search terms

1.	client education/
2.	health education/ or health knowledge/
3.	client participation/
4.	computer assisted instruction/
5.	Information Dissemination/ or Information Services/ or Information Seeking/ or Information/
6.	hot line services/
7.	social support/
8.	counseling/
9.	Support Groups/
10.	exp social networks/
11.	Peer Counseling/
12.	interpersonal communication/
13.	Communication Barriers/
14.	educational programs/
15.	client attitudes/
16.	health attitudes/
17.	client centered therapy/
18.	client satisfaction/
19.	or/1-18

A.2.5 Health economic (HE) search terms**Medline search terms**

1.	economics/
2.	value of life/
3.	exp "costs and cost analysis"/
4.	exp economics, hospital/
5.	exp economics, medical/
6.	economics, nursing/
7.	economics, pharmaceutical/
8.	exp "fees and charges"/
9.	exp budgets/
10.	budget*.ti,ab.
11.	cost*.ti.
12.	(economic* or pharmaco?economic*).ti.
13.	(price* or pricing*).ti,ab.
14.	(cost* adj2 (effective* or utilit* or benefit* or minimi* or unit* or estimat* or variable*)).ab.

15.	(financ* or fee or fees).ti,ab.
16.	(value adj2 (money or monetary)).ti,ab.
17.	or/1-16

Embase search terms

1.	health economics/
2.	exp economic evaluation/
3.	exp health care cost/
4.	exp fee/
5.	budget/
6.	funding/
7.	budget*.ti,ab.
8.	cost*.ti.
9.	(economic* or pharmaco?economic*).ti.
10.	(price* or pricing*).ti,ab.
11.	(cost* adj2 (effective* or utilit* or benefit* or minimi* or unit* or estimat* or variable*)).ab.
12.	(financ* or fee or fees).ti,ab.
13.	(value adj2 (money or monetary)).ti,ab.
14.	or/1-13

A.2.6 Quality of life (QOL) and model search terms**Medline search terms**

1.	quality-adjusted life years/
2.	sickness impact profile/
3.	(quality adj2 (wellbeing or well-being)).ti,ab.
4.	sickness impact profile.ti,ab.
5.	disability adjusted life.ti,ab.
6.	(qal* or qtime* or qwb* or daly*).ti,ab.
7.	(euroqol* or eq5d* or eq 5d*).ti,ab.
8.	(qol* or hql* or hqol* or h qol* or hrqol* or hr qol*).ti,ab.
9.	(health utility* or utility score* or disutilit*).ti,ab.
10.	(hui or hui1 or hui2 or hui3).ti,ab.
11.	health* year* equivalent*.ti,ab.
12.	(hye or hyes).ti,ab.
13.	rosser.ti,ab.
14.	(willingness to pay or time tradeoff or time trade off or tto or standard gamble*).ti,ab.
15.	(sf36 or sf 36 or short form 36 or shortform 36 or shortform36).ti,ab.
16.	(sf20 or sf 20 or short form 20 or shortform 20 or shortform20).ti,ab.
17.	(sf12 or sf 12 or short form 12 or shortform 12 or shortform12).ti,ab.
18.	(sf8 or sf 8 or short form 8 or shortform 8 or shortform8).ti,ab.
19.	(sf6 or sf 6 or short form 6 or shortform 6 or shortform6).ti,ab.
20.	Or/1-19

Embase search terms

1.	quality adjusted life year/
2.	"quality of life index"/

3.	short form 12/ or short form 20/ or short form 36/ or short form 8/
4.	sickness impact profile/
5.	(quality adj2 (wellbeing or well-being)).ti,ab.
6.	sickness impact profile.ti,ab.
7.	disability adjusted life.ti,ab.
8.	(qal* or qtime* or qwb* or daly*).ti,ab.
9.	(euroqol* or eq5d* or eq 5d*).ti,ab.
10.	(qol* or hq!* or hqol* or h qol* or hrqol* or hr qol*).ti,ab.
11.	(health utility* or utility score* or disutilit*).ti,ab.
12.	(hui or hui1 or hui2 or hui3).ti,ab.
13.	health* year* equivalent*.ti,ab.
14.	(hye or hyes).ti,ab.
15.	rosser.ti,ab.
16.	(willingness to pay or time tradeoff or time trade off or tto or standard gamble*).ti,ab.
17.	(sf36 or sf 36 or short form 36 or shortform 36 or shortform36).ti,ab.
18.	(sf20 or sf 20 or short form 20 or shortform 20 or shortform20).ti,ab.
19.	(sf12 or sf 12 or short form 12 or shortform 12 or shortform12).ti,ab.
20.	(sf8 or sf 8 or short form 8 or shortform 8 or shortform8).ti,ab.
21.	(sf6 or sf 6 or short form 6 or shortform 6 or shortform6).ti,ab.
22.	or/1-21

A.2.7 Excluded study designs and publication types search terms

The following study designs and publication types were removed from retrieved results using the NOT operator for **all** searches.

Medline search terms

1.	letter/
2.	editorial/
3.	news/
4.	exp historical article/
5.	anecdotes as topic/
6.	comment/
7.	case report/
8.	(letter or comment*).ti.
9.	or/1-8
10.	randomized controlled trial/ or random*.ti,ab.
11.	9 not 10
12.	animals/ not humans/
13.	exp animals, laboratory/
14.	exp animal experimentation/
15.	exp models, animal/
16.	exp rodentia/
17.	(rat or rats or mouse or mice).ti.
18.	or/11-17

Embase search terms

1.	letter.pt. or letter/
2.	note.pt.
3.	editorial.pt.
4.	case report/ or case study/
5.	(letter or comment*).ti.
6.	or/1-5
7.	randomized controlled trial/ or random*.ti,ab.
8.	6 not 7
9.	exp animal/ not human/
10.	nonhuman/
11.	exp experimental animal/
12.	exp animal experiment/
13.	exp animal model/
14.	exp rodent/
15.	(rat or rats or mouse or mice).ti.
16.	or/8-15

A.3 Searches by specific questions

A.3.1 Diagnostic criteria

What are the key diagnostic criteria for the following:

- Multiple sclerosis
- Possible multiple sclerosis
- Neuromyelitis optica
- Clinically isolated syndrome

Medline

Citation search of:

Polman, C. H., Reingold, S. C., Banwell, B., Clanet, M., Cohen, J. A., Filippi, M., Fujihara, K., Havrdova, E., Hutchinson, M., Kappos, L., Lublin, F. D., Montalban, X., O'Connor, P., Sandberg-Wollheim, M., Thompson, A. J., Waubant, E., Weinshenker, B. and Wolinsky, J. S. (2011), Diagnostic criteria for multiple sclerosis: 2010 Revisions to the McDonald criteria. *Ann Neurol.*, 69: 292–302. doi: 10.1002/ana.22366

A.3.2 Information and support

For adults with MS and their carers what information, education and support would they find useful?

Search constructed by combining the columns in the following table using the AND Boolean operator

Population	Intervention / exposure	Comparison	Study filter used	Date parameters
Adults who have a diagnosis of MS or possible MS, or are being investigated for MS. Including CIS	As below	n/a	The following filters were used in Medline, Embase and Psycinfo only: Patient	See Table 1

Population	Intervention / exposure	Comparison	Study filter used	Date parameters
			Information (PI)	

Medline search terms

1.	exp consumer health information/ or patient education as topic/
2.	patient education handout.pt.
3.	Access to Information/
4.	information centers/ or information services/
5.	Computer-Assisted Instruction/
6.	*Health Education/
7.	*Learning/
8.	publications/ or pamphlets/ or answering services/
9.	*communication/ or communication barriers/ or hotlines/ or information dissemination/
10.	social support/
11.	Health Communication/
12.	Self-Help Groups/
13.	*counseling/
14.	Patient Participation/
15.	Patient-Centered Care/
16.	"Patient Acceptance of Health Care"/
17.	attitude to health/ or health knowledge, attitudes, practice/
18.	((patient\$ or client\$ or carer\$ or caregiver\$) adj2 (helpline\$ or help line\$ or advice line\$ or newsletter\$ or website\$ or knowledge\$ or information or education or advoc\$ or literature or infopack\$ or leaflet\$ or pamphlet\$ or handout\$ or booklet\$ or message board\$ or workshop\$ or seminar\$ or discussion group\$ or brochure\$ or magazine\$ or factsheet\$ or counsel\$ or training)).ti,ab.
19.	((MS or multiple sclerosis) adj2 (resource center\$ or resource centre\$ or counselling or advisor or advisors or helpline\$ or help line\$ or advice line\$ or newsletter\$ or website\$ or infopack\$ or leaflet\$ or pamphlet\$ or brochure\$ or handout\$ or factsheet\$)).ti,ab.
20.	(need for information or information leaflet\$ or information needs or patient empowerment or peer support or patient support or emotional support or patient story or patient stories or access to information).ti,ab.
21.	((doctor\$ or physician\$ or clinician\$ or health professional\$ or nurse\$) adj2 (patient\$ or client\$ or carer\$ or caregiver\$) adj2 communic\$).ti,ab.
22.	(online adj2 (forum\$ or communiti\$)).ti,ab.
23.	((education\$ or information) adj2 (session\$ or provision or requirement\$ or support or program\$ or resource\$ or material\$ or intervention\$)).ti,ab.
24.	(support group\$ or support service\$ or self help group\$ or selfhelp group\$).ti,ab.
25.	(psychosocial adj2 (support or therap\$ or intervention\$)).ti,ab.
26.	((patient\$ or client\$ or carer\$ or caregiver\$) adj2 (perception\$ or preference\$ or expectation\$ or perspective\$ or view\$ or experience or experiences or belief\$ or opinion\$)).ti,ab.
27.	or/1-26

Embase search terms

1.	*patient education/
2.	*access to information/
3.	*information service/
4.	*information center/

5.	*health education/
6.	*patient information/
7.	*learning/
8.	*publication/
9.	*interpersonal communication/
10.	*information dissemination/
11.	*social support/
12.	*support group/
13.	*patient counseling/
14.	*patient attitude/ or *patient participation/
15.	*attitude to health/
16.	((patient\$ or client\$ or carer\$ or caregiver\$) adj2 (helpline\$ or help line\$ or advice line\$ or newsletter\$ or website\$ or knowledge\$ or information or education or advoc\$ or literature or infopack\$ or leaflet\$ or pamphlet\$ or handout\$ or booklet\$ or brochure\$ or message board\$ or workshop\$ or seminar\$ or discussion group\$ or magazine\$ or factsheet\$ or counsel\$ or training)).ti,ab.
17.	((MS or multiple sclerosis) adj2 (resource center\$ or resource centre\$ or counselling or advisor or advisors or helpline\$ or help line\$ or advice line\$ or newsletter\$ or website\$ or infopack\$ or leaflet\$ or pamphlet\$ or handout\$ or brochure\$ or factsheet\$)).ti,ab.
18.	(patient experienced outcomes or need for information or information leaflet\$ or information needs or patient empowerment or peer support or patient support or emotional support or patient story or patient stories or access to information).ti,ab.
19.	((doctor\$ or physician\$ or clinician\$ or health professional\$ or nurse\$) adj2 (patient\$ or carer\$ or caregiver\$ or client\$) adj2 communic\$).ti,ab.
20.	(online adj2 (forum\$ or communit\$)).ti,ab.
21.	((education\$ or information) adj2 (provision or requirement\$ or support or program\$ or resource\$ or session\$ or material\$ or intervention\$)).ti,ab.
22.	(support group\$ or support service\$ or self help group\$ or selfhelp group\$).ti,ab.
23.	(psychosocial adj2 (support or therap\$ or intervention\$)).ti,ab.
24.	((patient\$ or client\$ or carer\$ or caregiver\$) adj2 (perception\$ or preference\$ or expectation\$ or perspective\$ or view\$ or experience or experiences or belief\$ or opinion\$)).ti,ab.
25.	or/1-25

Cochrane search terms

#1.	MeSH descriptor: [Consumer Health Information] this term only
#2.	MeSH descriptor: [Patient Education as Topic] this term only
#3.	MeSH descriptor: [Access to Information] this term only
#4.	MeSH descriptor: [Information Centers] this term only
#5.	MeSH descriptor: [Information Services] this term only
#6.	MeSH descriptor: [Computer-Assisted Instruction] this term only
#7.	MeSH descriptor: [Health Education] this term only
#8.	MeSH descriptor: [Learning] this term only
#9.	MeSH descriptor: [Publications] this term only
#10.	MeSH descriptor: [Pamphlets] this term only
#11.	MeSH descriptor: [Answering Services] this term only
#12.	MeSH descriptor: [Communication] this term only
#13.	MeSH descriptor: [Hotlines] this term only
#14.	MeSH descriptor: [Communication Barriers] this term only

#15.	MeSH descriptor: [Information Dissemination] this term only
#16.	MeSH descriptor: [Social Support] this term only
#17.	MeSH descriptor: [Health Communication] this term only
#18.	MeSH descriptor: [Self-Help Groups] this term only
#19.	MeSH descriptor: [Counseling] this term only
#20.	MeSH descriptor: [Patient Participation] this term only
#21.	MeSH descriptor: [Patient-Centered Care] this term only
#22.	MeSH descriptor: [Patient Acceptance of Health Care] this term only
#23.	MeSH descriptor: [Attitude to Health] this term only
#24.	MeSH descriptor: [Health Knowledge, Attitudes, Practice] this term only
#25.	((patient* or client* or carer* or caregiver*) near/2 (helpline* or "help line" or "help lines" or "advice line" or "advice lines" or newsletter* or website* or knowledge or information or "education" or advoc* or literature)):ti,ab
#26.	((patient* or client* or carer* or caregiver*) near/2 (infopack* or leaflet* or pamphlet* or handout* or booklet* or "message board" or "message boards" or workshop* or seminar* or "discussion group" or "discussion groups" or brochure* or magazine* or factsheet* or counsel* or training)):ti,ab
#27.	((MS or "multiple sclerosis") near/2 ("resource center" or "resource centers" or "resource centre" or "resource centres" or counselling or advisors or "advisor" or helpline* or "help line" or "help lines" or "advice line" or "advice lines" or newsletter* or website* or infopack* or leaflet* or pamphlet* or brochure* or handout* or factsheet*)):ti,ab
#28.	("need for information" or "patient experienced outcomes" or "information leaflet" or "information leaflets" or "information needs" or "patient empowerment" or "peer support" or "patient support" or "emotional support" or "patient story" or "patient stories" or "access to information"):ti,ab
#29.	((doctor* or nurs* or clinician* or physician* or "health professional" or "health professionals") near/2 (patient* or client* or carer or caregiver)) near/2 communicat*):ti,ab
#30.	(online near/2 (forum* or communit*)):ti,ab
#31.	((education* or information) near/2 (session* or provision or requirement* or support or program* or resource* or material* or intervention*)):ti,ab
#32.	((support or selfhelp or "self help") near/2 (group* or service*)):ti,ab
#33.	(psychosocial near/2 (support or therap* or intervention*)):ti,ab
#34.	((patient* or client* or carer* or caregiver*) near/2 (perception* or preferenc* or expectation* or perspectiv* or view* or "experience" or "experiences" or belief* or opinion*)):ti,ab
#35.	MeSH descriptor: [Patient Preference] this term only
#36.	{or#1-#35}

Psycinfo search terms

1.	((patient or client\$ or carer\$ or caregiver\$) adj2 (helpline\$ or help line\$ or advice line\$ or newsletter\$ or website\$ or knowledge\$ or information or education or advoc\$ or literature or infopack\$ or leaflet\$ or pamphlet\$ or handout\$ or factsheet\$ or counsel\$ or training)).ti,ab.
2.	((patient or client\$ or carer\$ or caregiver\$) adj2 (booklet\$ or brochure\$ or message board\$ or workshop\$ or seminar\$ or discussion group\$ or magazine\$)).ti,ab.
3.	((MS or multiple sclerosis) adj2 (resource center\$ or resource centre\$ or counselling or advisor or advisors or helpline\$ or help line\$ or advice line\$ or newsletter\$ or website\$ or infopack\$ or leaflet\$ or pamphlet\$ or handout\$ or factsheet\$)).ti,ab.
4.	(patient experienced outcomes or need for information or information leaflet\$ or information needs or patient empowerment or peer support or patient support or emotional support or patient story or patient stories or access to information).ti,ab.

5.	((doctor\$ or physician\$ or clinician\$ or nurs\$ or health professional\$) adj2 (patient\$ or client\$ or carer\$ or caregiver\$) adj2 communic\$).ti,ab.
6.	(online adj2 (forum\$ or communit\$)).ti,ab.
7.	((education\$ or information) adj2 (session\$ or provision or requirement\$ or support or program\$ or resource\$ or material\$ or intervention\$)).ti,ab.
8.	(support group\$ or support service\$ or self help group\$ or selfhelp group\$).ti,ab.
9.	(psychosocial adj2 (support or therap\$ or intervention\$)).ti,ab.
10.	((patient\$ or client\$ or carer\$ or caregiver\$) adj2 (perception\$ or preference\$ or expectation\$ or perspective\$ or view\$ or experience or experiences or belief\$ or opinion\$)).ti,ab.
11.	or/1-10

A.3.3 Modifiable risk factors

For adults with MS what advice should be given to regarding modifiable risk factors such as immunisations and pregnancy?

Search constructed by combining the columns in the following table using the AND Boolean operator

Population	Intervention / exposure	Comparison	Study filter used	Date parameters
Adults who have a diagnosis of MS or possible MS, or are being investigated for MS.	As below	n/a	The following filters were used in all databases OBS	See Table 1

Medline search terms

1.	exp Immunization/
2.	(immuniz* or vaccinat* or innoculat*).ti,ab.
3.	*Pregnancy/
4.	(childfree or "child free" or childless or pregnant or pregnancy).ti,ab.
5.	exp Smoking/
6.	(smoking or smoker or tobbaço or cigarette).ti,ab.
7.	*Stress, Psychological/
8.	(suffering adj2 (mental or psychologic* or lfe or emotion* or mental)).ti,ab.
9.	(stress adj2 (disorder* or psychologic* or life or emotion* or mental)).ti,ab.
10.	(anguish or anxiety or anxious or worried or worries or fear*).ti,ab.
11.	((stressful or life) adj2 event*).ti,ab.
12.	*exercise/
13.	*physical fitness/
14.	*resistance training/
15.	exp *sports/
16.	*leisure activities/
17.	*recreation/
18.	(physical adj3 (exercis\$ or exertion or endurance or therap\$ or conditioning or activit\$ or fitness)).ti,ab.
19.	(exercise adj3 (train\$ or intervention\$ or protocol\$ or program\$ or therap\$ or activit\$ or regim\$)).ti,ab.
20.	(fitness adj3 (train\$ or intervention\$ or protocol\$ or program\$ or therap\$ or activit\$ or regim\$ or centre\$ or center\$)).ti,ab.

21.	((training or conditioning) adj3 (intervention\$ or protocol\$ or program\$ or activit\$ or regim\$)).ti,ab.
22.	(sport\$ or recreation\$ or leisure or cycling or bicycl\$ or rowing or treadmill\$ or running or circuit training or swim\$ or walk\$ or dance\$ or dancing or tai ji or tai chi or taiji or taichi or yoga).ti,ab.
23.	((endurance or aerobic or cardio\$) adj3 (fitness or train\$ or intervention\$ or protocol\$ or program\$ or therap\$ or activit\$ or regim\$)).ti,ab.
24.	(muscle strengthening or progressive resist\$).ti,ab.
25.	((weight or strength\$ or resistance) adj (train\$ or lift\$ or exercise\$)).ti,ab.
26.	((isometric or isotonic or eccentric or concentric) adj (action\$ or contraction\$ or exercise\$)).ti,ab.
27.	or/1-26

Embase search terms

1.	exp Immunization/
2.	(immuniz* or vaccinat* or innoculat*).ti,ab.
3.	*Pregnancy/
4.	(childfree or "child free" or childless or pregnant or pregnancy).ti,ab.
5.	exp Smoking/
6.	(smoking or smoker or tobbaço or cigarette).ti,ab.
7.	*Mental stress/
8.	(suffering adj2 (mental or psychologic* or lfe or emotion* or mental)).ti,ab.
9.	(stress adj2 (disorder* or psychologic* or life or emotion* or mental)).ti,ab.
10.	(anguish or anxiety or anxious or worried or worries or fear*).ti,ab.
11.	((stressful or life) adj2 event*).ti,ab.
12.	*exercise/ or *aerobic exercise/
13.	*arm exercise/ or *athletic performance/ or *dynamic exercise/ or *exercise intensity/ or *isokinetic exercise/ or *muscle exercise/ or *pilates/ or *static exercise/
14.	*locomotion/ or *swimming/ or *walking/ or *dancing/ or *physical activity/ or *jumping/ or *lifting effort/ or *stretching/ or *weight lifting/ or *fitness/
15.	(physical adj3 (exercis\$ or exertion or endurance or therap\$ or conditioning or activit\$ or fitness)).ti,ab.
16.	(exercise adj3 (train\$ or intervention\$ or protocol\$ or program\$ or therap\$ or activit\$ or regim\$)).ti,ab.
17.	(fitness adj3 (train\$ or intervention\$ or protocol\$ or program\$ or therap\$ or activit\$ or regim\$ or centre\$ or center\$)).ti,ab.
18.	((training or conditioning) adj3 (intervention\$ or protocol\$ or program\$ or activit\$ or regim\$)).ti,ab.
19.	(sport\$ or recreation\$ or leisure or cycling or bicycl\$ or rowing or treadmill\$ or running or circuit training or swim\$ or walk\$ or dance\$ or dancing or tai ji or tai chi or taiji or taichi or yoga).ti,ab.
20.	((endurance or aerobic or cardio\$) adj3 (fitness or train\$ or intervention\$ or protocol\$ or program\$ or therap\$ or activit\$ or regim\$)).ti,ab.
21.	(muscle strengthening or progressive resist\$).ti,ab.
22.	((weight or strength\$ or resistance) adj (train\$ or lift\$ or exercise\$)).ti,ab.
23.	((isometric or isotonic or eccentric or concentric) adj (action\$ or contraction\$ or exercise\$)).ti,ab.
24.	or/1-24

Cochrane search terms

#1.	MeSH descriptor: [Immunization] explode all trees
#2.	MeSH descriptor: [Smoking] explode all trees
#3.	MeSH descriptor: [Pregnancy] explode all trees
#4.	MeSH descriptor: [Stress, Psychological] explode all trees
#5.	MeSH descriptor: [Exercise] explode all trees
#6.	{or #1-#5}

A.3.4 Structured assessment

Searches for the following three questions were run as one search:

Does the use of structural assessment(s) compared with non-structured assessment(s) improve patient and carer outcomes for young people and adults with MS for the following disabilities and functional problems?

What is the optimal timing of a structure assessment?

What should be the frequency of a structured assessment?

Search constructed by combining the columns in the following table using the AND Boolean operator

Population	Intervention / exposure	Comparison	Study filter used	Date parameters
Adults who have a diagnosis of MS or possible MS, or are being investigated for MS. Including CIS.	As below	n/a	The following filters were used in Medline and Embase only: RCT, SR	See Table 1

Medline search terms

1.	Neurologic Examination/
2.	Multiple Sclerosis/cl
3.	Disability Evaluation/
4.	((disability or comprehensive or clinical or structured) adj2 (evaluation* or assessment* or review* or battery)).ti,ab.
5.	((regular* or routine or consistent or recur* or repeat* or frequent*) adj2 (evaluation* or assessment* or review* or battery)).ti,ab.
6.	((Disability or neurologic* or functional or Walking or Impact or Performance) adj3 (Scale or assessment or Composite)).ti,ab.
7.	(Kurtzke* or DSS or GNDS or EDSS or HAQUAMS or Hamburg Quality of Life or MSIS 29 or MSIS29 or MSQI or MSQOL 54 or MSQOL54 or MSWS 12 or MSWS 12 or PS MS or PSMS or Qual OT or QualOT or SI MS or Symptom Inventory or FAMS or NFI or CAMBS or MSFC or Leeds MS or Cambridge multiple sclerosis).ti,ab.
8.	(Quality of Life in Occupational Therapy or Quality of Life Inventory).ti,ab.
9.	or/1-8

Embase search terms

1.	neurologic examination/ or neuropsychological test/
2.	disability severity/
3.	"named inventories, questionnaires and rating scales"/ or expanded disability status scale/
4.	((disability or comprehensive or clinical or structured) adj2 (evaluation* or assessment* or

	review* or battery)).ti,ab.
5.	((regular* or routine or consistent or recur* or repeat* or frequent*) adj2 (evaluation* or assessment* or review* or battery)).ti,ab.
6.	((Disability or neurologic* or functional or Walking or Impact or Performance) adj3 (Scale or assessment or Composite)).ti,ab.
7.	(Kurtzke* or DSS or GNDS or EDSS or HAQUAMS or Hamburg Quality of Life or MSIS 29 or MSIS29 or MSQI or MSQOL 54 or MSQOL54 or MSWS 12 or MSWS 12 or PS MS or PSMS or Qual OT or QualOT or SI MS or Symptom Inventory or FAMS or NFI or CAMBS or MSFC or Leeds MS or Cambridge multiple sclerosis).ti,ab.
8.	(Quality of Life in Occupational Therapy or Quality of Life Inventory).ti,ab.
9.	or/1-8

Cochrane search terms

#1.	MeSH descriptor: [Neurologic Examination] this term only
#2.	MeSH descriptor: [Multiple Sclerosis] this term only and with qualifier(s): [Classification - CL]
#3.	MeSH descriptor: [Disability Evaluation] this term only
#4.	((disability or comprehensive or clinical or structured) near/2 (evaluation* or assessment* or review* or battery)):ti,ab,kw
#5.	((regular* or routine or consistent or recur* or repeat* or frequent*) near/2 (evaluation* or assessment* or review* or battery)):ti,ab,kw
#6.	((Disability or neurologic* or functional or Walking or Impact or Performance) near/3 (Scale or assessment or Composite)):ti,ab,kw
#7.	(Kurtzke* or DSS or GNDS or EDSS or HAQUAMS or Hamburg Quality of Life or MSIS 29 or MSIS29 or MSQI or MSQOL 54 or MSQOL54 or MSWS 12 or MSWS 12 or PS MS or PSMS or Qual OT or QualOT or SI MS or Symptom Inventory or FAMS or NFI or CAMBS or MSFC or Leeds MS or Cambridge multiple sclerosis):ti,ab,kw
#8.	(Quality of Life in Occupational Therapy or Quality of Life Inventory):ti,ab,kw
#9.	#1 or #2 or #3 or #4 or #5 or #6 or #7 or #8

A.3.5 Coordination of care

For adults with MS and their carers, what interventions improve continuity of care?

Search constructed by combining the columns in the following table using the AND Boolean operator

Population	Intervention / exposure	Comparison	Study filter used	Date parameters
Adults who have a diagnosis of MS or possible MS, or are being investigated for MS.	As below	n/a	No study filters used	See Table 1

Medline search terms

1.	exp *Multiple Sclerosis/nu [Nursing]
2.	nursing evaluation research/ or nursing methodology research/
3.	Nurse's Role/
4.	Nurse Clinicians/
5.	models, nursing/
6.	Specialties, Nursing/
7.	patient care team/ or nursing, team/
8.	"Continuity of Patient Care"/

9.	exp *Medical Records Systems, Computerized/
10.	*patient care planning/ or critical pathways/
11.	*patient care management/ or *"quality of health care"/ or "process assessment (health care)"/
12.	*Comprehensive Health Care/
13.	"delivery of health care, integrated"/ or *health services accessibility/
14.	exp *Interprofessional Relations/
15.	*Program Evaluation/
16.	*Cooperative Behavior/
17.	*Medical Record Linkage/
18.	*Program Development/
19.	*models, organizational/
20.	*Case Management/
21.	((specialist\$ or MS) adj2 nurse\$).ti,ab.
22.	((continuity or continuum or co-ordinat\$) adj2 care).ti,ab.
23.	integrated care pathway\$.ti,ab.
24.	(multidisciplinary adj2 (team\$ or teamwork or care)).ti,ab.
25.	(model\$ adj2 care).ti,ab.
26.	(team-based adj2 (approach or treatment or care)).ti,ab.
27.	(key-worker or key worker or keyworker).ti,ab.
28.	((handover or hand-over) adj2 (routine\$ or procedure\$ or practice\$)).ti,ab.
29.	((regular\$ or proactiv\$) adj2 (review\$ or monitor\$ or check-up\$ or checkup\$ or followup\$ or follow-up\$)).ti,ab.
30.	((electronic or central\$) adj2 patient record\$).ti,ab.
31.	or/1-29

Embase search terms

1.	((specialist\$ or MS) adj2 nurse\$).ti,ab.
2.	((continuity or continuum or co-ordinat\$) adj2 care).ti,ab.
3.	integrated care pathway\$.ti,ab.
4.	(multidisciplinary adj2 (team\$ or teamwork or care)).ti,ab.
5.	(model\$ adj2 care).ti,ab.
6.	(team-based adj2 (approach or treatment or care)).ti,ab.
7.	(key-worker or key worker or keyworker).ti,ab.
8.	((handover or hand-over) adj2 (routine\$ or procedure\$ or practice\$)).ti,ab.
9.	((regular\$ or proactiv\$) adj2 (review\$ or monitor\$ or check-up\$ or checkup\$ or followup\$ or follow-up\$)).ti,ab.
10.	((electronic or central\$) adj2 patient record\$).ti,ab.
11.	*nurse/ or *nurse attitude/ or *nursing/ or *nursing care/
12.	*nurse patient relationship/
13.	nursing methodology research/
14.	nursing evaluation research/
15.	clinical nurse specialist/
16.	*nursing discipline/ or *nursing process/
17.	*patient care/ or *patient care planning/
18.	*health care quality/ or *"quality of nursing care"/

19.	*clinical pathway/
20.	*electronic medical record/
21.	*patient monitoring/ or *periodic medical examination/
22.	*integrated health care system/
23.	*interdisciplinary communication/
24.	*cooperation/
25.	*program development/
26.	*case management/
27.	*process model/
28.	or/1-27

Cochrane search terms

#1.	MeSH descriptor: [Multiple Sclerosis] explode all trees and with qualifier(s): [Nursing - NU]
#2.	MeSH descriptor: [Nursing Evaluation Research] this term only
#3.	MeSH descriptor: [Nursing Methodology Research] this term only
#4.	MeSH descriptor: [Nurse's Role] this term only
#5.	MeSH descriptor: [Nurse Clinicians] this term only
#6.	MeSH descriptor: [Models, Nursing] this term only
#7.	MeSH descriptor: [Specialties, Nursing] this term only
#8.	MeSH descriptor: [Patient Care Team] this term only
#9.	MeSH descriptor: [Nursing, Team] this term only
#10.	MeSH descriptor: [Continuity of Patient Care] this term only
#11.	MeSH descriptor: [Medical Records Systems, Computerized] explode all trees
#12.	MeSH descriptor: [Patient Care Planning] this term only
#13.	MeSH descriptor: [Critical Pathways] this term only
#14.	MeSH descriptor: [Patient Care Management] this term only
#15.	MeSH descriptor: [Quality of Health Care] this term only
#16.	MeSH descriptor: [Process Assessment (Health Care)] this term only
#17.	MeSH descriptor: [Comprehensive Health Care] this term only
#18.	MeSH descriptor: [Delivery of Health Care, Integrated] this term only
#19.	MeSH descriptor: [Interprofessional Relations] explode all trees
#20.	MeSH descriptor: [Program Evaluation] this term only
#21.	MeSH descriptor: [Cooperative Behavior] this term only
#22.	MeSH descriptor: [Medical Record Linkage] this term only
#23.	MeSH descriptor: [Program Development] this term only
#24.	MeSH descriptor: [Models, Organizational] this term only
#25.	MeSH descriptor: [Case Management] this term only
#26.	((specialist* or MS) near/2 nurse*):ti,ab
#27.	((continuity or continuum or co-ordinat*) near/2 care):ti,ab
#28.	((("integrated care") next/1 pathway*):ti,ab
#29.	((multi-disciplinary or multidisciplinary) next/2 (team* or teamwork or care)):ti,ab
#30.	(model next/2 care):ti,ab
#31.	(team-based near/2 (approach or treatment or care)):ti,ab
#32.	(key-worker or "key worker" or keyworker):ti,ab
#33.	((handover or hand-over) near/2 (routine* or procedure* or practice*)):ti,ab

#34.	((regular or proactiv*) near/2 (review* or monitor* or check-up* or checkup* or followup* or follow-up*)):ti,ab
#35.	((electronic or central*) near/2 ("patient records")):ti,ab
#36.	{or #1-#35}

A.3.6 Management of acute relapses

For adults with MS, what is the clinical evidence and cost effectiveness of pharmacological management of acute relapse with steroids?

Search constructed by combining the columns in the following table using the AND Boolean operator

Population	Intervention / exposure	Comparison	Study filter used	Date parameters
Adults who have a diagnosis of MS or possible MS, or are being investigated for MS.	As below	n/a	The following filters were used in Medline and Embase only: RCT, SR	See Table 1

Medline search terms

1.	adrenal cortex hormones/ or exp glucocorticoids/ or *steroids/
2.	(steroid\$ or corticoid\$ or glucocorticoid\$ or corticosteroid\$).ti,ab.
3.	(prednisolone or methylprednisolone or prednisone or dexamethasone or depo-medrone or solu-medrone or medrone or dexsol or martapan or deltacortril).ti,ab.
4.	1 or 2 or 3

Embase search terms

1.	*corticosteroid/
2.	exp *glucocorticoid/
3.	*steroid/
4.	steroid therapy/ or corticosteroid therapy/
5.	(prednisolone or methylprednisolone or prednisone or dexamethasone or depo-medron or solu-medrone or medrone or dexsol or martapan or deltacortril).ti,ab.
6.	(steroid\$ or corticoid\$ or glucocorticoid\$ or corticosteroid\$).ti,ab.
7.	or/1-6

Cochrane search terms

#1.	MeSH descriptor: [Adrenal Cortex Hormones] this term only
#2.	MeSH descriptor: [Glucocorticoids] explode all trees
#3.	MeSH descriptor: [Steroids] this term only
#4.	(solu-medrone or medrone or depo-medrone or methylprednisolone or prednisolone or prednisone or dexamethasone or dexsol or martapan or deltacortril or corticosteroid* or steroid*):ti,ab,kw
#5.	(glucocorticoid* or corticoid*):ti,ab,kw
#6.	#1 or #2 or #3 or #4 or #5

A.3.7 Pharmacological treatment of spasticity

For adults with MS, what is the clinical evidence and cost effectiveness of pharmacological treatment of spasticity?

Search constructed by combining the columns in the following table using the AND Boolean operator

Population	Intervention / exposure	Comparison	Study filter used	Date parameters
No population terms used	As below	n/a	The following filters were used in Medline and Embase only: RCT, SR	See Table 1

Medline search terms

1.	Baclofen/
2.	Baclofen.ti,ab.
3.	1 or 2

Embase search terms

1.	Baclofen/
2.	Baclofen.ti,ab.
3.	1 or 2

Cochrane search terms

#1.	MeSH descriptor: [Baclofen] explode all trees
#2.	(baclofen*):ti,ab
#3.	#1 or #2

A.3.8 Pharmacological treatment of mobility

For adults with MS, what is the clinical evidence and cost effectiveness of pharmacological treatment of mobility with fampridine compared to usual care?

Search constructed by combining the columns in the following table using the AND Boolean operator

Population	Intervention / exposure	Comparison	Study filter used	Date parameters
Adults who have a diagnosis of MS or possible MS, or are being investigated for MS. Including CIS.	As below	n/a	No study filters used	See Table 1

Medline search terms

1.	fampridine.mp.
2.	fampyra.ti,ab.
3.	Aminopyridine derivative/
4.	4-aminopyridine/
5.	Aminopyridine*.ti,ab.
6.	or/1-5

Embase search terms

1.	fampridine.mp.
2.	fampyra.ti,ab.
3.	Aminopyridine derivative/

4.	4-aminopyridine/
5.	Aminopyridine*.ti,ab.
6.	or/1-5

Cochrane search terms

#1.	MeSH descriptor Aminopyridines explode all trees
#2.	MeSH descriptor 4-Aminopyridine explode all trees
#3.	(fampridine):ti,ab,kw
#4.	(fampyra):ti,ab,kw
#5.	(Aminopyridine*):ti,ab,kw
#6.	(#1 OR #2 OR #3 OR #4 OR #5)

A.3.9 Pharmacological treatment of fatigue

For adults with MS, what is the clinical evidence and cost effectiveness of pharmacological treatment of fatigue (amantadine, B12 injections, SSRIs)?

Search constructed by combining the columns in the following table using the AND Boolean operator

Population	Intervention / exposure	Comparison	Study filter used	Date parameters
Adults who have a diagnosis of MS or possible MS, or are being investigated for MS.	As below	n/a	The following filters were used in Medline and Embase only: RCT, SR	See Table 1

Medline search terms

1.	Aspirin/
2.	(Aspirin or Acetylsalicylic acid or ASA or Acetysal or Acylpyrin or Aloxiprimum or Colfarit or Dispril or Easprin or Ecotrin or Endosprin or Magnecyl or Micristin or Polopirin or Polopirynga or Solprin or Solupsan or Zorprin or Acetyloxy benzoic Acid).ti,ab.
3.	exp Amantadine/
4.	(Amantadin* or Lysovir or Symmetrel or Aman or merz or ratiopharm or neuraxpharm or amantahciazu or amanta or mantadix or cerebramed or amantadinratiopharm or tregor or midantan or pmsamantadine or wiregyt or "1 aminoadamantane" or ratiopharm or azupharma or amantasulfateazu or adamantylamine or endantadine or symadine or genamantadine or amixx or adekin or viregyt or gen-amantadine or infex).ti,ab.
5.	exp Serotonin Uptake Inhibitors/ or exp Citalopram/ or exp Paroxetine/ or exp Fluoxetine/
6.	(SSRI* or Citalopram or Cytalopram or Cipramil or Cipralex or Lexapro or escitalopram or Fluoxetine or Prozac or oxactin or prozit or Fluvoxamine or Faverin or sarafem or Paroxetine or Seroxat or Sertraline or Lustral or paxil or arropax).ti,ab.
7.	((serotonin or inhibitor) adj2 (uptake or reuptake)).ti,ab.
8.	exp Vitamin B 12/ or exp Hydroxocobalamin/
9.	((B12 adj1 vitamin) or cyanocobalamin or cobalamin or eritron or cobalamins or cytamen or hydroxycobalamin).ti,ab.
10.	or/1-9

Embase search terms

	acetylsalicylic acid/
1.	(Aspirin or Acetylsalicylic acid or ASA or Acetysal or Acylpyrin or Aloxiprimum or Colfarit or Dispril or Easprin or Ecotrin or Endosprin or Magnecyl or Micristin or Polopirin or Polopirynga or

	Solprin or Solupsan or Zorprin or Acetyloxy benzoic Acid).ti,ab.
2.	exp amantadine derivative/ or exp amantadine/ or exp amantadine sulfate/
3.	(Amantadin* or Lysovir or Symmetrel or Aman or merz or ratiopharm or neuraxpharm or amantahciazu or amanta or mantadix or cerebramed or amantadinratiopharm or tregor or midantan or pmsamantadine or wiregyt or "1 aminoadamantane" or ratiopharm or azupharma or amantasulfateazu or adamantylamine or endantadine or symadine or genamantadine or amixx or adekin or viregyt or gen-amantadine or infex).ti,ab.
4.	exp serotonin uptake inhibitor/
5.	(SSRI* or Citalopram or Cytalopram or Cipramil or CipraleX or Lexapro or escitalopram or Fluoxetine or Prozac or oxactin or prozit or Fluvoxamine or Faverin or sarafem or Paroxetine or Seroxat or Sertraline or Lustral or paxil or aropax).ti,ab.
6.	((serotonin or inhibitor) adj2 (uptake or reuptake)).ti,ab.
7.	exp cyanocobalamin/
8.	exp hydroxocobalamin/
9.	((B12 adj1 vitamin) or cyanocobalamin or cobalamin or eritron or cobalamins or cytamen or hydroxycobalamin).ti,ab.
10.	or/1-9

Cochrane search terms

#1.	MeSH descriptor: [Aspirin] this term only
#2.	(Aspirin or Acetylsalicylic acid or ASA or Acetysal or Acylpyrin or AloxiPrimum or Colfarit or Dispril or Easprin or Ecotrin or Endosprin or Magnecyl or Micristin or Polopirin or Polopiryna or Solprin or Solupsan or Zorprin or Acetyloxy benzoic Acid):ti,ab
#3.	MeSH descriptor: [Amantadine] this term only
#4.	(Amantadin* or Lysovir or Symmetrel or Aman or merz or ratiopharm or neuraxpharm or amantahciazu or amanta or mantadix or cerebramed or amantadinratiopharm or tregor or midantan or pmsamantadine or wiregyt or "1 aminoadamantane" or ratiopharm or azupharma or amantasulfateazu or adamantylamine or endantadine or symadine or genamantadine or amixx or adekin or viregyt or gen-amantadine or infex):ti,ab
#5.	Serotonin near/2 inhibitor:ti,ab
#6.	uptake near/2 inhibitor*:ti,ab
#7.	reuptake near/2 inhibitor*:ti,ab
#8.	(SSRI* or Citalopram or Cytalopram or Cipramil or ESCITALOPRAM or CipraleX or Lexapro or Fluoxetine or Prozac or oxactin or prozit or Fluvoxamine or Faverin or sarafem or Paroxetine or Seroxat or Sertraline or Lustral or paxil or aropax):ti,ab
#9.	MeSH descriptor: [Serotonin Uptake Inhibitors] explode all trees
#10.	MeSH descriptor: [Vitamin B 12] explode all trees
#11.	MeSH descriptor: [Hydroxocobalamin] explode all trees
#12.	("B12 vitamin" or "vitamin B12" or cyanocobalamin or cobalamin or eritron or cobalamins or cytamen or hydroxycobalamin):ti,ab
#13.	{or #1-#13}

A.3.10 Pharmacological treatment of oscillopsia

For the following population a different population was used as outlined below.

For adults with MS, what is the clinical evidence and cost effectiveness of pharmacological treatment of oscillopsia?

Search constructed by combining the columns in the following table using the AND Boolean operator

Population	Intervention / exposure	Comparison	Study filter used	Date parameters
Adults who have a diagnosis of MS or possible MS, or are being investigated for MS and oscillopsia or nystagmus. (Population terms adapted as below)	As below	n/a	No study filters used	See Table 1

Medline search terms

1.	(oscillopsia or nystagmus).ti,ab.
2.	((ocular or eye\$) adj3 (movement\$ or oscillat\$)).ti,ab.
3.	Ocular Motility Disorders/
4.	or/1-3
5.	[MS POP from section A.1.1. AND 4]
6.	exp Nystagmus, Physiologic/ or Nystagmus, Pathologic/
7.	(acquired adj4 nystagmus).ti,ab.
8.	[5 or 6 or 7] Population for this question
9.	Baclofen/
10.	(baclofen or lioresal).ti,ab.
11.	gamma-Aminobutyric Acid/
12.	(gabapentin or fanatrex or gabarone or neogab or gralise or neurontin or nupentin).ti,ab.
13.	Memantine/
14.	(memantine or ebixa).ti,ab.
15.	Piracetam/
16.	(levetiracetam or keppra).ti,ab.
17.	exp Botulinum Toxins/
18.	(botulinum* or onabotulinum* or rimabotulinum* or abobotulinum*).ti,ab.
19.	(dysport or botox or btx or oculinum or vistabel or azzalure or bocouture or xeomin or reloxin or prosigne or purtox or nt201 or mybloc or neurobloc).ti,ab.
20.	Clonazepam/
21.	(clonazepam or rivotril).ti,ab.
22.	Isoniazid/
23.	isoniazid.ti,ab.
24.	Valproic Acid/
25.	(valproate or depakote or convulex).ti,ab.
26.	muscarinic antagonists/ or benzotropine/ or scopolamine hydrobromide/ or trihexyphenidyl/
27.	(benztropine or benztropine or cogentin or scopolamine or hyoscine or scopoderm or trihexyphenidyl or broflex or artane).ti,ab.
28.	(antimuscarinic\$ or anti-muscarinic).ti,ab.
29.	(muscarinic adj4 (antagonist\$ or blocker\$ or blocking agent\$)).ti,ab.
30.	gaba agonists/ or gaba-b receptor agonists/
31.	Cholinergic Antagonists/
32.	((anti-cholinergic or anticholinergic) adj3 (drug\$ or agent\$)).ti,ab.

33.	(cholinergic adj4 (antagonist\$ or blocker\$ or blocking agent\$)).ti,ab.
34.	or/9-33
35.	8 and 34

Embase search terms

1.	oscillopsia/
2.	eye movement disorder/
3.	(oscillopsia or nystagmus).ti,ab.
4.	((ocular or eye\$) adj3 (movement\$ or oscillat\$)).ti,ab.
5.	or/1-4
6.	[MS POP from section A.1.1. AND 5]
7.	*nystagmus/ or *optokinetic nystagmus/ or *"pendular nystagmus"/
8.	(acquired adj4 nystagmus).ti,ab.
9.	[6 or 7 or 8] Population for this question
10.	*baclofen/
11.	(baclofen or lioresal).ti,ab.
12.	*gabapentin/
13.	(gabapentin or fanatrex or gabarone or neogab or gralise or neurontin or nupentin).ti,ab.
14.	*memantine/
15.	(memantine or ebixa).ti,ab.
16.	*etiracetam/
17.	(levetiracetam or keppra).ti,ab.
18.	*botulinum toxin B/
19.	*botulinum toxin A/ or *botulinum toxin/
20.	(botulinum* or onabotulinum* or rimabotulinum* or abobotulinum*).ti,ab.
21.	(dysport or botox or btx or oculinum or xeomin or reloxin or vistabel or azzalure or bocouture or prosigne or purtox or nt201 or mybloc or neurobloc).ti,ab.
22.	*clonazepam/
23.	(clonazepam or rivotril).ti,ab.
24.	*isoniazid/ or *isoniazid derivative/
25.	isoniazid.ti,ab.
26.	*valproic acid/
27.	(valproate or depakote or convulex).ti,ab.
28.	*muscarinic receptor blocking agent/ or *scopolamine/ or *trihexyphenidyl/
29.	*benzatropine/
30.	(benztropine or benztropine or scopolamine or scopoderm ot hyoscine or trihexyphenidyl or broflex or artane).ti,ab.
31.	(antimuscarinic\$ or anti-muscarinic).ti,ab.
32.	(muscarinic adj4 (antagonist\$ or blocking agent\$ or blocker\$)).ti,ab.
33.	*4 aminobutyric acid receptor stimulating agent/
34.	*scopolamine bromide/ or *scopolamine butyl bromide/
35.	*cholinergic receptor blocking agent/
36.	((anti-cholinergic or anticholinergic) adj3 (agent\$ or drug\$)).ti,ab.
37.	(cholinergic adj4 (antagonist\$ or blocking agent\$ or blocker\$)).ti,ab.
38.	or/10-37
39.	9 and 38

Cochrane search terms

#1.	(oscillopsia or nystagmus):ti,ab,kw
#2.	((ocular or eye*) next/3 (movement* or oscillat*)):ti,ab
#3.	MeSH descriptor: [Ocular Motility Disorders] this term only
#4.	#1 or #2 or #3
#5.	[MS POP from section A.1.1. and #4]
#6.	MeSH descriptor: [Nystagmus, Pathologic] this term only
#7.	MeSH descriptor: [Nystagmus, Physiologic] explode all trees
#8.	(acquired near/4 nystagmus):ti,ab
#9.	[#5 or #6 or #7 or #8] Population for this question
#10.	(gabapentin or baclofen or lioresal or memantine or fanatrex or gabarone or neogab or gralise or neurontin or nupentin or memantine or ebixa or levetiracetam or keppra or botulinam* or onabotulinum* or rimabotulinum* or abobotulinum* or dysport or botox or btx or oculinum or vistabel or azzalure or bocouture or xeomin or relaxin or prosigne or purtox or nt201 or mybloc or neurobloc or clonazepam or rivotril or isoniazid or valproate or depakote or convulex):ti,ab,kw
#11.	(benztropine or benzatropine or cogentin or scopolamine or hyoscine or scopoderm or trihexyphenidyl or broflex or artane):ti,ab,kw
#12.	((antimuscarinic or anti-muscarinic or anticholinergic or anti-cholinergic) near/3 (drug* or agent*)):ti,ab
#13.	((muscarinic or cholinergic) near/3 (antagonist* or blocker* or "blocking agent" or "blocking agents")):ti,ab
#14.	MeSH descriptor: [Baclofen] this term only
#15.	MeSH descriptor: [gamma-Aminobutyric Acid] this term only
#16.	MeSH descriptor: [Memantine] this term only
#17.	MeSH descriptor: [Piracetam] this term only
#18.	MeSH descriptor: [Botulinum Toxins] explode all trees
#19.	MeSH descriptor: [Clonazepam] this term only
#20.	MeSH descriptor: [Isoniazid] this term only
#21.	MeSH descriptor: [Valproic Acid] this term only
#22.	MeSH descriptor: [Muscarinic Antagonists] this term only
#23.	MeSH descriptor: [Benztropine] this term only
#24.	MeSH descriptor: [Scopolamine Hydrobromide] explode all trees
#25.	MeSH descriptor: [Trihexyphenidyl] this term only
#26.	MeSH descriptor: [Cholinergic Antagonists] this term only
#27.	MeSH descriptor: [GABA Agonists] this term only
#28.	MeSH descriptor: [Receptors, GABA-B] this term only
#29.	#10 or #11 or #12 or #13 or #14 or #15 or #16 or #17 or #18 or #19 or #20 or #21 or #22 or #23 or #24 or #25 or #26 or #27 or #28
#30.	#9 and #29

A.3.11 Pharmacological treatment of emotionalism

For adults with MS, what is the clinical evidence and cost effectiveness of pharmacological management of emotionalism with antidepressants?

Search constructed by combining the columns in the following table using the AND Boolean operator

Population	Intervention / exposure	Comparison	Study filter used	Date parameters
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Population	Intervention / exposure	Comparison	Study filter used	Date parameters
Adults who have a diagnosis of MS or possible MS, or are being investigated for MS.	As below	n/a	The following filters were used in Medline, Embase and Psycinfo only: RCT, SR	See Table 1

Medline search terms

1.	exp Antidepressive Agents/
2.	(Antidepress* or anti depress*).ab,ti.
3.	exp *Serotonin Uptake Inhibitors/
4.	SSRI*.ti,ab.
5.	(serotonin adj3 (uptake or re-uptake or reuptake or re uptake)).ti,ab.
6.	Citalopram/ or Fluoxetine/ or Fluvoxamine/ or Paroxetine/ or Sertraline/
7.	(citalopram or cipramil or Escitalopram or cipralex or fluoxetine or prozac or fluvoxamine or faverin or paroxetine or seroxat or sertraline or lustral).ti,ab.
8.	(SNRI* or SSNRI*).ti,ab.
9.	Flupentixol/ or tryptophan/
10.	(duloxetine or cymbalta or yentreve or Venlafaxine or efexor or agomelatine or valdoxan or flupentixol or fluvoxol or depixel or reboxetine or edronanx or tryptophan or optimax).ti,ab.
11.	NaSSA*.ti,ab.
12.	Mianserin/ or Trazodone/
13.	(Mianserin or Mirtazapine or zipirin or Trazodone or molipaxin).ti,ab.
14.	exp *antidepressive agents, tricyclic/
15.	(tricyclic* or TCAs).ti,ab.
16.	Amitriptyline/ or Clomipramine/ or Dothiepin/ or Doxepin/ or Imipramine/ or Lofepamine/ or Nortriptyline/ or Trimipramine/
17.	(amitriptyline or triptafen or Clomipramine or Anafranil or Dothiepin or Dosulepin or Prothiaden or teofarma or Doxepin or sinepin or Imipramine or Lofepamine or Nortriptyline or Allegron or Trimipramine or Surmontil).ti,ab.
18.	exp *Monoamine Oxidase Inhibitors/
19.	(Monoamine Oxidase Inhibitor* or MAOI*).ti,ab.
20.	Phenelzine/ or Isocarboxazid/ or Tranylcypromine/ or Moclobemide/
21.	(Phenelzine or nardil or Isocarboxazid or Tranylcypromine or Moclobemide or manerix).ti,ab.
22.	Dextromethorphan/
23.	Dextromethorphan.ti,ab.
24.	*Antipsychotic Agents/
25.	((("second generation" or "2nd Generation" or atypical) adj2 (anti psychotic* or antipsychotic*)).ti,ab.
26.	AAPs.ti,ab.
27.	Clozapine/ or Risperidone/
28.	(Amisulpride or Solian or Aripiprazole or Abilify or Clozapine or clozaril or denzapine or zaponex or Olanzapine or zyprexa or Paliperidone or invega or Quetiapine or seroquel or Risperidone or risperdal).ti,ab.
29.	*Anticonvulsants/
30.	(anti adj2 (epileptic* or convulsant* or seizure*)).ti,ab.

31.	(antiepileptic* or anticonvulsant* or AEDs or antiseizure).ti,ab.
32.	Carbamazepine/ or Ethosuximide/ or Phenobarbital/ or Primidone/ or Phenytoin/ or Valproic Acid/ or Vigabatrin/ or Benzodiazepines/ or Clonazepam/ or Diazepam/ or Lorazepam/ or Midazolam/ or Phenobarbital/ or Phenytoin/ or Acetazolamide/ or Piracetam/
33.	(Carbamazepine or tegretol or Eslicarbazepine or Zebinix or Oxcarbazepine or Trileptal or Ethosuximide or emeside or zarontin or Gabapentin or neurotin or Pregabalin or Lyrica or Lacosamide or vimpat or Lamotrigine or Lamictal or Levetiracetam or keppra or Perampanel or fycopa or Phenobarbital or Phenobarbitone or Primidone or Mysoline or Phenytoin or epanutin or Retigabine or trobalt or Rufinamide or inovelon or Tiagabine or gabitril or Topiramate or topamax or sodium valproate or epilim or episenta or epival or valproic acid or convulex or depakote or vigabatrin or sabril or Zonisamide or zonegran or Benzodiazepine* or Clobazam or Clonazepam or rivotril or Diazepam or Fosphenytoin or pro-epanutin or Lorazepam or Midazolam or buccolam or Phenobarbital or Phenytoin or Epanutin or Acetazolamide or diamox or Piracetam or nootropil).ti,ab.
34.	or/1-33

Embase search terms

1.	exp Antidepressive Agent/
2.	(Antidepress* or anti depress*).ab,ti.
3.	exp *Serotonin Uptake Inhibitor/
4.	SSRI*.ti,ab.
5.	(serotonin adj3 (uptake or re-uptake or reuptake or re uptake)).ti,ab.
6.	*Citalopram/ or *Escitalopram/ or *Fluoxetine/ or *Fluvoxamine/ or *Paroxetine/ or *Sertraline/
7.	(citalopram or cipramil or Escitalopram or cipralex or fluoxetine or prozac or fluvoxamine or faverin or paroxetine or seroxat or sertraline or lustral).ti,ab.
8.	exp *serotonin noradrenalin reuptake inhibitor/
9.	(SNRI* or SSNRI*).ti,ab.
10.	*Duloxetine/ or *Venlafaxine/ or *agomelatine/ or *flupentixol/ or *reboxetine/ or *tryptophan/
11.	(duloxetine or cymbalta or yentreve or Venlafaxine or efexor or agomelatine or valdoxan or flupentixol or fluanxol or depixol or reboxetine or edronanx or tryptophan or optimax).ti,ab.
12.	NaSSA*.ti,ab.
13.	*Mianserin/ or *Mirtazapine/ or *Trazodone/
14.	(Mianserin or Mirtazapine or zipirin or trazodone or molipaxin).ti,ab.
15.	exp *tricyclic antidepressant agent/
16.	(tricyclic* or TCAs).ti,ab.
17.	*Amitriptyline/ or *Clomipramine/ or *Dothiepin/ or *Doxepin/ or *Imipramine/ or *Lofepamine/ or *Nortriptyline/ or *Trimipramine/
18.	(amitriptyline or triptafen or Clomipramine or Anafranil or Dothiepin or Dosulepin or Prothiaden or teofarma or Doxepin or sinepin or Imipramine or Lofepamine or Nortriptyline or Allegron or Trimipramine or Surmontil).ti,ab.
19.	exp *Monoamine Oxidase Inhibitor/
20.	(Monoamine Oxidase Inhibitor* or MAOI*).ti,ab.
21.	*Phenelzine/ or *Isocarboxazid/ or *Tranlycypromine/ or *Moclobemide/
22.	(Phenelzine or nardil or Isocarboxazid or Tranlycypromine or Moclobemide or manerix).ti,ab.
23.	*Dextromethorphan/
24.	Dextromethorphan.ti,ab.
25.	exp *neuroleptic agent/

26.	((("second generation" or "2nd Generation" or atypical) adj2 (anti psychotic* or antipsychotic*)).ti,ab.
27.	AAPs.ti,ab.
28.	*Amisulpride/ or *Aripiprazole/ or *Clozapine/ or *Olanzapine/ or *Paliperidone/ or *Quetiapine/ or *Risperidone/
29.	(Amisulpride or Solian or Aripiprazole or Abilify or Clozapine or clozaril or denzapine or zaponex or Olanzapine or zyprexa or Paliperidone or invega or Quetiapine or seroquel or Risperidone or risperdal).ti,ab.
30.	exp *anticonvulsive agent/
31.	(anti adj2 (epileptic* or convulsant* or seizure*)).ti,ab.
32.	(antiepileptic* or anticonvulsant* or AEDs or antiseizure).ti,ab.
33.	*Carbamazepine/ or *Eslicarbazepine/ or *Oxcarbazepine/ or *Ethosuximide/ or *Gabapentin/ or *Pregabalin/ or *Lacosamide/ or *Lamotrigine/ or *Levetiracetam/ or *Levetiracetam/ or *Perampanel/ or *Phenobarbital/ or *Primidone/ or *Phenytoin/ or *Retigabine/ or *Rufinamide/ or *Tiagabine/ or *Topiramate/ or *Valproic Acid/ or *Vigabatrin/ or *Zonisamide/ or *Benzodiazepines/ or *Clobazam/ or *Clonazepam/ or *Diazepam/ or *Fosphenytoin/ or *Lorazepam/ or *Midazolam/ or *Phenobarbital/ or *Phenytoin/ or *Acetazolamide/ or *Piracetam/
34.	(Carbamazepine or tegretol or Eslicarbazepine or Zebinix or Oxcarbazepine or Trileptal or Ethosuximide or emeside or zarontin or Gabapentin or neurotin or Pregabalin or Lyrica or Lacosamide or vimpat or Lamotrigine or Lamictal or Levetiracetam or keppra or Perampanel or fycompa or Phenobarbital or Phenobarbitone or Primidone or mysoline or Phenytoin or epanutin or Retigabine or trobalt or Rufinamide or inovelon or Tiagabine or gabitril or Topiramate or topamax or sodium valproate or epilim or episenta or epival or valproic acid or convulex or depakote or vigabatrin or sabril or Zonisamide or zonegran or Benzodiazepine* or Clobazam or Clonazepam or rivotril or Diazepam or Fosphenytoin or pro-epanutin or Lorazepam or Midazolam or buccolam or Phenobarbital or Phenytoin or Epanutin or Acetazolamide or diamox or Piracetam or nootropil).ti,ab.
35.	or/1-34

Cochrane search terms

#1.	MeSH descriptor: [Serotonin Uptake Inhibitors] this term only
#2.	SSRI*:ti,ab
#3.	(serotonin near/3 (uptake or re-uptake or reuptake or re uptake)):ti,ab
#4.	MeSH descriptor: [Citalopram] explode all trees
#5.	MeSH descriptor: [Fluoxetine] explode all trees
#6.	MeSH descriptor: [Fluvoxamine] explode all trees
#7.	MeSH descriptor: [Paroxetine] explode all trees
#8.	MeSH descriptor: [Sertraline] explode all trees
#9.	(citalopram or cipramil or Escitalopram or cipralext or fluoxetine or prozac or fluvoxamine or faverin or paroxetine or seroxat or sertraline or lustral):ti,ab
#10.	(SNRI* or SSNRI*):ti,ab
#11.	MeSH descriptor: [Flupenthixol] explode all trees
#12.	MeSH descriptor: [Tryptophan] explode all trees
#13.	(duloxetine or cymbalta or yentreve or Venlafaxine or efexor or agomelatine or valdoxan or flupenthixol or fluaxnol or depixel or reboxetine or edronanx or tryptophan or optimax):ti,ab
#14.	NaSSA*:ti,ab
#15.	MeSH descriptor: [Mianserin] explode all trees
#16.	MeSH descriptor: [Trazodone] explode all trees
#17.	(Mianserin or Mirtazapine or zipirin or Trazodone or molipaxin):ti,ab

#18.	MeSH descriptor: [Antidepressive Agents, Tricyclic] explode all trees
#19.	(tricyclic* or TCAs):ti,ab
#20.	MeSH descriptor: [Amitriptyline] explode all trees
#21.	MeSH descriptor: [Clomipramine] explode all trees
#22.	MeSH descriptor: [Dothiepin] explode all trees
#23.	MeSH descriptor: [Doxepin] explode all trees
#24.	MeSH descriptor: [Imipramine] explode all trees
#25.	MeSH descriptor: [Lofepramine] explode all trees
#26.	MeSH descriptor: [Nortriptyline] explode all trees
#27.	MeSH descriptor: [Trimipramine] explode all trees
#28.	(amitriptyline or triptafen or Clomipramine or Anafranil or Dothiepin or Dosulepin or Prothiaden or teofarma or Doxepin or sinepin or Imipramine or Lofepramine or Nortriptyline or Allegron or Trimipramine or Surmontil):ti,ab
#29.	MeSH descriptor: [Monoamine Oxidase Inhibitors] explode all trees
#30.	(Monoamine Oxidase Inhibitor* or MAOI*):ti,ab
#31.	MeSH descriptor: [Phenelzine] explode all trees
#32.	MeSH descriptor: [Isocarboxazid] explode all trees
#33.	MeSH descriptor: [Tranlycypromine] explode all trees
#34.	MeSH descriptor: [Moclobemide] explode all trees
#35.	(Phenelzine or nardil or Isocarboxazid or Tranlycypromine or Moclobemide or manerix):ti,ab
#36.	MeSH descriptor: [Dextromethorphan] explode all trees
#37.	Dextromethorphan:ti,ab
#38.	MeSH descriptor: [Antipsychotic Agents] explode all trees
#39.	((("second generation" or "2nd Generation" or atypical) near/2 (anti psychotic* or antipsychotic*)):ti,ab (Word variations have been searched)
#40.	AAPs:ti,ab
#41.	MeSH descriptor: [Clozapine] explode all trees
#42.	MeSH descriptor: [Risperidone] explode all trees
#43.	(Amisulpride or Solian or Aripiprazole or Abilify or Clozapine or clozaril or denzapine or zaponex or Olanzapine or zyprexa or Paliperidone or invega or Quetiapine or seroquel or Risperidone or risperdal):ti,ab
#44.	MeSH descriptor: [Anticonvulsants] explode all trees
#45.	(anti near/2 (epileptic* or convulsant* or seizure*)):ti,ab
#46.	(antiepileptic* or anticonvulsant* or AEDs or antiseizure):ti,ab
#47.	MeSH descriptor: [Carbamazepine] explode all trees
#48.	MeSH descriptor: [Ethosuximide] explode all trees
#49.	MeSH descriptor: [Phenobarbital] explode all trees
#50.	MeSH descriptor: [Primidone] explode all trees
#51.	MeSH descriptor: [Phenytoin] explode all trees
#52.	MeSH descriptor: [Valproic Acid] explode all trees
#53.	MeSH descriptor: [Vigabatrin] explode all trees
#54.	MeSH descriptor: [Benzodiazepines] explode all trees
#55.	MeSH descriptor: [Clonazepam] explode all trees
#56.	MeSH descriptor: [Diazepam] explode all trees
#57.	MeSH descriptor: [Lorazepam] explode all trees
#58.	MeSH descriptor: [Midazolam] explode all trees

#59.	MeSH descriptor: [Phenobarbital] explode all trees
#60.	MeSH descriptor: [Phenytoin] explode all trees
#61.	MeSH descriptor: [Acetazolamide] explode all trees
#62.	MeSH descriptor: [Piracetam] explode all trees
#63.	(Carbamazepine or tegretol or Eslicarbazepine or Zebinix or Oxcarbazepine or Trileptal or Ethosuximide or emeside or zarontin or Gabapentin or neurotin or Pregabalin or Lyrica or Lacosamide or vimpat or Lamotrigine or Lamictal or Levetiracetam or keppra or Pirampanel or fycompa or Phenobarbital or Phenobarbitone or Primidone or mysoline or Phenytoin or epanutin or Retigabine or trobalt or Rufinamide or inovelon or Tiagabine or gabitril or Topiramate or topamax or sodium valproate or epilim or episenta or epival or valproic acid or convulex or depakote or vigabatrin or sabril or Zonisamide or zonegran or Benzodiazepine* or Clobazam or Clonazepam or rivotril or Diazepam or Fosphenytoin or pro-epanutin or Lorazepam or Midazolam or buccolam or Phenobarbital or Phenytoin or Epanutin or Acetazolamide or diamox or Piracetam or nootropil):ti,ab
#64.	{or #1-#63}

Psycinfo search terms

1.	exp Serotonin Reuptake Inhibitors/
2.	SSRI*.ti,ab.
3.	(serotonin adj3 (uptake or re-uptake or reuptake or re uptake)).ti,ab.
4.	Citalopram/ or Fluoxetine/ or Fluvoxamine/ or Paroxetine/ or Sertraline/
5.	(citalopram or cipramil or Escitalopram or cipralex or fluoxetine or prozac or fluvoxamine or faverin or paroxetine or seroxat or sertraline or lustral).ti,ab.
6.	(SNRI* or SSNRI*).ti,ab.
7.	Flupentixol/ or tryptophan/
8.	(duloxetine or cymbalta or yentreve or Venlafaxine or efexor or agomelatine or valdoxan or flupentixol or fluaxol or depixel or reboxetine or edronanx or tryptophan or optimax).ti,ab.
9.	NaSSA*.ti,ab.
10.	Mianserin/ or Trazodone/
11.	(Mianserin or Mirtazapine or zipirin or Trazodone or molipaxin).ti,ab.
12.	exp Antidepressant Drugs/
13.	(tricyclic* or TCAs).ti,ab.
14.	Amitriptyline/ or Clomipramine/ or Dothiepin/ or Doxepin/ or Imipramine/ or Lofepramine/ or Nortriptyline/ or Trimipramine/
15.	(amitriptyline or triptafen or Clomipramine or Anafranil or Dothiepin or Dosulepin or Prothiaden or teofarma or Doxepin or sinepin or Imipramine or Lofepramine or Nortriptyline or Allegron or Trimipramine or Surmontil).ti,ab.
16.	exp *Monoamine Oxidase Inhibitors/
17.	(Monoamine Oxidase Inhibitor* or MAOI*).ti,ab.
18.	Phenelzine/ or Isocarboxazid/ or Tranylcypromine/ or Moclobemide/
19.	(Phenelzine or nardil or Isocarboxazid or Tranylcypromine or Moclobemide or manerix).ti,ab.
20.	Dextromethorphan.ti,ab.
21.	Neuroleptic drugs/
22.	((("second generation" or "2nd Generation" or atypical) adj2 (anti psychotic* or antipsychotic*)).ti,ab.
23.	AAPs.ti,ab.
24.	Clozapine/ or Risperidone/
25.	(Amisulpride or Solian or Aripiprazole or Abilify or Clozapine or clozaril or denzapine or zaponex or Olanzapine or zyprexa or Paliperidone or invega or Quetiapine or seroquel or

	Risperidone or risperdal).ti,ab.
26.	exp *Anticonvulsive Drugs/
27.	(anti adj2 (epileptic* or convulsant* or seizure*).ti,ab.
28.	(antiepileptic* or anticonvulsant* or AEDs or antiseizure).ti,ab.
29.	Carbamazepine/ or Ethosuximide/ or Phenobarbital/ or Primidone/ or Phenytoin/ or Valproic Acid/ or Vigabatrin/ or Benzodiazepines/ or Clonazepam/ or Diazepam/ or Lorazepam/ or Midazolam/ or Phenobarbital/ or Phenytoin/ or Acetazolamide/ or Piracetam/
30.	(Carbamazepine or tegretol or Eslicarbazepine or Zebinix or Oxcarbazepine or Trileptal or Ethosuximide or emeside or zarontin or Gabapentin or neurotin or Pregabalin or Lyrica or Lacosamide or vimpat or Lamotrigine or Lamictal or Levetiracetam or keppra or Perampanel or fycompa or Phenobarbital or Phenobarbitone or Primidone or mysoline or Phenytoin or epanutin or Retigabine or trobalt or Rufinamide or inovelon or Tiagabine or gabitril or Topiramate or topamax or sodium valproate or epilim or episenta or epival or valproic acid or convulex or depakote or vigabatrin or sabril or Zonisamide or zonegran or Benzodiazepine* or Clobazam or Clonazepam or rivotril or Diazepam or Fosphenytoin or pro-epanutin or Lorazepam or Midazolam or buccolam or Phenobarbital or Phenytoin or Epanutin or Acetazolamide or diamox or Piracetam or nootropil).ti,ab.
31.	or/1-30

A.3.12 Pharmacological treatment of ataxia and tremor

For adults with MS, what is the clinical evidence and cost effectiveness of pharmacological treatment of ataxia and tremor with baclofen?

Search constructed by combining the columns in the following table using the AND Boolean operator

Population	Intervention / exposure	Comparison	Study filter used	Date parameters
Adults who have a diagnosis of MS or possible MS, or are being investigated for MS.	As below	n/a	The following filters were used in Medline and Embase only: RCT, SR	See Table 1

Medline search terms

1.	Baclofen/ or aminopyridines/ or *4-aminopyridine/
2.	(baclofen* or baclophen or lioresal or kemstro or fampridine or fampyra or aminopyridine*).ti,ab.
3.	exp Isoniazid/
4.	Isoniazid.ti,ab.
5.	*Anticonvulsants/
6.	(anti adj2 (epileptic* or convulsant* or seizure*).ti,ab.
7.	(antiepileptic* or anticonvulsant* or AEDs or antiseizure).ti,ab.
8.	Carbamazepine/ or Ethosuximide/ or Phenobarbital/ or Primidone/ or Phenytoin/ or Valproic Acid/ or Vigabatrin/ or Benzodiazepines/ or Clonazepam/ or Diazepam/ or Lorazepam/ or Midazolam/ or Phenobarbital/ or Phenytoin/ or Acetazolamide/ or Piracetam/
9.	(Carbamazepine or tegretol or Eslicarbazepine or Zebinix or Oxcarbazepine or Trileptal or Ethosuximide or emeside or zarontin or Gabapentin or neurotin or Pregabalin or Lyrica or Lacosamide or vimpat or Lamotrigine or Lamictal or Levetiracetam or keppra or Perampanel or fycompa or Phenobarbital or Phenobarbitone or Primidone or mysoline or Phenytoin or epanutin or Retigabine or trobalt or Rufinamide or inovelon or Tiagabine or gabitril or Topiramate or topamax or sodium valproate or epilim or episenta or epival or valproic acid or convulex or depakote or vigabatrin or sabril or Zonisamide or zonegran or Benzodiazepine* or Clobazam or Clonazepam or rivotril or Diazepam or Fosphenytoin or pro-epanutin or

	Lorazepam or Midazolam or buccolam or Phenobarbital or Phenytoin or Epanutin or Acetazolamide or diamox or Piracetam or nootropil).ti,ab.
10.	exp Adrenergic beta-Antagonists/
11.	(propranolol or angilol or angilol or inderal-la or half-inderal or inderal or bedranol or prograne or slo-pro or acebutolol or sectral or atenolol or tenormin or bisoprolol or cardicor or emcor or carvedilol or eucardic or celiprolol or celectol or co-tenidone or tenoret or tenoretic or esmolol or brevibloc or labetalol or trandate or metoprolol or betaloc or lopresor or nadolol or corgard or nebivolol or nebilet or hypoloc or oxprenolol or trasicor or slow-trasicor or pindolol or visken or sotalol or beta-cardone or sotacor or timolol or betim).ti,ab.
12.	(beta adj2 antagonist*).ti,ab.
13.	((b or beta) adj3 block*).ti,ab.
14.	((beta-adrenoceptor or b-adrenoceptor or beta-adrenergic) adj3 (blockade or blocker* or blocking or antagonist*).ti,ab.
15.	Ondansetron/
16.	(ondansetron or zofran).ti,ab.
17.	exp Botulinum Toxins/
18.	(botulinum adj3 (a or b)).ti,ab.
19.	(dysport or botox or btx or oculinum or xeomin or relaxin or prosigne or purtox or nt201 or mybloc or neurobloc).ti,ab.
20.	Cannabis/
21.	Cannabinoids/
22.	(Cannabis or cannabinoids or marihuana or marijuana or ganja or hasish or sativex).ti,ab.
23.	Dantrolene/ or Methocarbamol/ or Tramadol/ or Temazepam/ or Flunitrazepam/ or Flurazepam/ or Nitrazepam/
24.	(dantrolene or dantrium or tizanidine or zanaflex or methocarbamol or robaxin or tramadol or zamadol or zydol or larapam or mabron or marol or maxitram or tramquel or zeridame or tradorec or nitrazepam or temazepam or flunitrazepam or dalmene or zaleplon or sonata or zolpidem or stilnoct or zoiclone or zimovane).ti,ab.
25.	fampridine.mp.
26.	fampyra.ti,ab.
27.	aminopyridines/ or *4-aminopyridine/
28.	Aminopyridine*.ti,ab.
29.	or/1-28

Embase search terms

1.	Baclofen/ or Aminopyridine derivative/ or *4-aminopyridine/
2.	(baclofen* or baclophen or lioesal or kemstro or fampridine or fampyra or aminopyridine*).ti,ab.
3.	exp Isoniazid/
4.	Isoniazid.ti,ab.
5.	exp *anticonvulsive agent/
6.	(anti adj2 (epileptic* or convulsant* or seizure*).ti,ab.
7.	(antiepileptic* or anticonvulsant* or AEDs or antiseizure).ti,ab.
8.	*Carbamazepine/ or *Eslicarbazepine/ or *Oxcarbazepine/ or *Ethosuximide/ or *Gabapentin/ or *Pregabalin/ or *Lacosamide/ or *Lamotrigine/ or *Levetiracetam/ or *Levetiracetam/ or *Perampanel/ or *Phenobarbital/ or *Primidone/ or *Phenytoin/ or *Retigabine/ or *Rufinamide/ or *Tiagabine/ or *Topiramate/ or *Valproic Acid/ or *Vigabatrin/ or *Zonisamide/ or *Benzodiazepines/ or *Clobazam/ or *Clonazepam/ or *Diazepam/ or *Fosphenytoin/ or *Lorazepam/ or *Midazolam/ or *Phenobarbital/ or *Phenytoin/ or *Acetazolamide/ or *Piracetam/

9.	(Carbamazepine or tegretol or Eslicarbazepine or Zebinix or Oxcarbazepine or Trileptal or Ethosuximide or emeside or zarontin or Gabapentin or neurotin or Pregabalin or Lyrica or Lacosamide or vimpat or Lamotrigine or Lamictal or Levetiracetam or keppra or Perampanel or fycampa or Phenobarbital or Phenobarbitone or Primidone or mysoline or Phenytoin or epanutin or Retigabine or trobalt or Rufinamide or inovelon or Tiagabine or gabitril or Topiramate or topamax or sodium valproate or epilim or episenta or epival or valproic acid or convulex or depakote or vigabatrin or sabril or Zonisamide or zonegran or Benzodiazepine* or Clobazam or Clonazepam or rivotril or Diazepam or Fosphenytoin or pro-epanutin or Lorazepam or Midazolam or buccolam or Phenobarbital or Phenytoin or Epanutin or Acetazolamide or diamox or Piracetam or nootropil).ti,ab.
10.	exp *beta adrenergic receptor blocking agent/
11.	exp *bisoprolol/ or exp *bisoprolol fumarate/ or exp *bisoprolol fumarate plus hydrochlorothiazide/ or exp *carvedilol/ or exp *metoprolol/ or exp*metoprolol fumarate/ or exp *metoprolol succinate/ or exp *metoprolol tartrate/ or exp *nebivolol/
12.	(propranolol or angilol or angilol or inderal-1a or half-inalderal or inderal or bedranol or prograne or slo-pro or acebutolol or sectral or atenolol or tenormin or bisoprolol or cardicor or emcor or carvedilol or eucardic or celiprolol or celectol or co-tenidone or tenoret or tenoretic or esmolol or brevilbloc or labetalol or trandate or metoprolol or betaloc or lopresor or nadolol or corgard or nebivolol or nebilet or hypoloc or oxprenolol or trasicor or slow-trasicor or pindolol or visken or sotalol or beta-cardone or sotacor or timolol or betim).ti,ab.
13.	(beta adj2 antagonist*).ti,ab.
14.	((b or beta) adj3 block*).ti,ab.
15.	((beta-adrenoceptor or b-adrenoceptor or beta-adrenergic) adj3 (blockade or blocker* or blocking or antagonist*).ti,ab.
16.	Ondansetron/
17.	(ondansetron or zofran).ti,ab.
18.	exp botulinum toxin/ or exp botulinum toxin b/ or exp botulinum toxin a/
19.	(botulinum adj3 (a or b)).ti,ab.
20.	(dysport or botox or btx or oculinum or xeomin or relaxin or prosigne or purtox or nt201 or mybloc or neurobloc).ti,ab.
21.	Cannabis/
22.	Cannabinoid/
23.	(Cannabis or cannabinoids or marihuana or marijuana or ganja or hasish or sativex).ti,ab.
24.	Dantrolene/ or Methocarbamol/ or Tramadol/ or Temazepam/ or Flunitrazepam/ or Flurazepam/ or Nitrazepam/
25.	(dantrolene or dantrium or tizandine or zanaflex or methocarbamol or robaxin or tramadol or zamadol or zydol or larapam or mabron or marol or maxitram or tramquel or zeridame or tradorec or nitrazepam or temazepam or flunitrazepam or dalmane or zaleplon or sonata or zolpidem or stilnoct or zoiclone or zimovane).ti,ab.
26.	fampridine.mp.
27.	fampyra.ti,ab.
28.	Aminopyridine derivative/
29.	4-aminopyridine/
30.	Aminopyridine*.ti,ab.
31.	or/1-30

Cochrane search terms

#1.	MeSH descriptor: [Aminopyridines] explode all trees
#2.	MeSH descriptor: [4-Aminopyridine] explode all trees
#3.	fampridine:ti,ab,kw

#4.	fampyra:ti,ab,kw
#5.	Aminopyridine*:ti,ab,kw
#6.	MeSH descriptor: [Baclofen] this term only
#7.	(baclofen* or baclophen or lioesal or kemstro):ti,ab,kw
#8.	MeSH descriptor: [Adrenergic beta-Antagonists] explode all trees
#9.	beta-blocker*:ti
#10.	(propranolol or angilol or angilol or inderal-1a or half-inalderal or inderal or bedranol or prograne or slo-pro or acebutolol or sectral or atenolol or tenormin or bisoprolol or cardicor or emcor or carvedilol or eucardic or celiprolol or celectol or co-tenidone or tenoret or tenoretic or esmolol or brevilbloc or labetalol or trandate or metoprolol or betaloc or lopresor or nadolol or corgard or nebivolol or nebilet or hypoloc or oxprenolol or trasicor or slow-trasicor or pindolol or visken or sotalol or beta-cardone or sotacor or timolol or betim):ti,ab,kw
#11.	MeSH descriptor: [Anticonvulsants] explode all trees
#12.	(anti near/2 (epileptic* or convulsant* or seizure*)):ti,ab
#13.	(antiepileptic* or anticonvulsant* or AEDs or antiseizure):ti,ab
#14.	MeSH descriptor: [Carbamazepine] explode all trees
#15.	MeSH descriptor: [Ethosuximide] explode all trees
#16.	MeSH descriptor: [Phenobarbital] explode all trees
#17.	MeSH descriptor: [Primidone] explode all trees
#18.	MeSH descriptor: [Phenytoin] explode all trees
#19.	MeSH descriptor: [Valproic Acid] explode all trees
#20.	MeSH descriptor: [Vigabatrin] explode all trees
#21.	MeSH descriptor: [Benzodiazepines] explode all trees
#22.	MeSH descriptor: [Clonazepam] explode all trees
#23.	MeSH descriptor: [Diazepam] explode all trees
#24.	MeSH descriptor: [Lorazepam] explode all trees
#25.	MeSH descriptor: [Midazolam] explode all trees
#26.	MeSH descriptor: [Phenobarbital] explode all trees
#27.	MeSH descriptor: [Phenytoin] explode all trees
#28.	MeSH descriptor: [Acetazolamide] explode all trees
#29.	MeSH descriptor: [Piracetam] explode all trees
#30.	(Carbamazepine or tegretol or Eslicarbazepine or Zebinix or Oxcarbazepine or Trileptal or Ethosuximide or emeside or zarontin or Gabapentin or neurotin or Pregabalin or Lyrica or Lacosamide or vimpat or Lamotrigine or Lamictal or Levetiracetam or keppra or Perampanel or fycompa or Phenobarbital or Phenobarbitone or Primidone or mysoline or Phenytoin or epanutin or Retigabine or trobalt or Rufinamide or inovelon or Tiagabine or gabitril or Topiramate or topamax or sodium valproate or epilim or episenta or epival or valproic acid or convulex or depakote or vigabatrin or sabril or Zonisamide or zonegran or Benzodiazepine* or Clobazam or Clonazepam or rivotril or Diazepam or Fosphenytoin or pro-epanutin or Lorazepam or Midazolam or buccolam or Phenobarbital or Phenytoin or Epanutin or Acetazolamide or diamox or Piracetam or nootropil):ti,ab
#31.	MeSH descriptor: [Cannabis] explode all trees
#32.	MeSH descriptor: [Isoniazid] explode all trees
#33.	Isoniazid:ti,ab,kw
#34.	MeSH descriptor: [Ondansetron] explode all trees
#35.	(ondansetron or zofran):ti,ab,kw
#36.	MeSH descriptor: [Botulinum Toxins] explode all trees
#37.	(botulinum near/3 (a or b)):ti,ab,kw

#38.	(dysport or botox or btx or oculinum or xeomin or relaxin or prosigne or purtox or nt201 or mybloc or neurobloc):ti,ab
#39.	MeSH descriptor: [Cannabis] explode all trees
#40.	MeSH descriptor: [Cannabinoids] explode all trees
#41.	(Cannabis or cannabinoids or marihuana or marijuana or ganja or hasish or sativex) .ti,ab.
#42.	MeSH descriptor: [Dantrolene] explode all trees
#43.	MeSH descriptor: [Methocarbamol] explode all trees
#44.	MeSH descriptor: [Tramadol] explode all trees
#45.	MeSH descriptor: [Temazepam] explode all trees
#46.	MeSH descriptor: [Flunitrazepam] explode all trees
#47.	MeSH descriptor: [Flurazepam] explode all trees
#48.	MeSH descriptor: [Nitrazepam] explode all trees
#49.	(dantrolene or dantrium or tizanidine or zanaflex or methocarbamol or robaxin or tramadol or zamadol or zydol or larapam or mabron or marol or maxitram or tramquel or zeridame or tradorec or nitrazepam or temazepam or flunitrazepam or dalmene or zaleplon or sonata or zolpidem or stilnoct or zoiclone or zimovane):ti,ab
#50.	{or #1-#49}

A.3.13 Non pharmacological treatment of memory and cognition

For adults with MS, what is the clinical evidence and cost effectiveness of non-pharmacological management of memory and cognitive problems with neuropsychological rehabilitation?

Search constructed by combining the columns in the following table using the AND Boolean operator

Population	Intervention / exposure	Comparison	Study filter used	Date parameters
Adults who have a diagnosis of MS or possible MS, or are being investigated for MS.	As below	n/a	The following filters were used in Medline, Embase and PsycINFO only: RCT, SR	See Table 1

Medline search terms

1.	exp Memory/ or exp Memory Disorders/ or exp Cognition/ or exp cognition disorders/ or Attention/ or Extinction, Psychological/ or auditory perceptual disorders/
2.	neuropsychology/ or language/ or exp speech/ or problem solving/ or mathematics/ or exp learning/ or exp thinking/ or exp psycholinguistics/
3.	((perceptual or attentional) adj3 (disorders* or deficit* or impairment* or abilit*)).ti,ab.
4.	(cogniti* or neuropsychol* or memor* or learn* or attenti* or information process* or language or visuopat* or visuocognit* or problem solving or reason or execut* or metacognit*).ti,ab.
5.	or/1-4
6.	Rehabilitation/ or therapeutics/ or therapy, computer-assisted/ or exp Neuropsychological Tests/
7.	Remedial teaching/ or "recovery of function"/ or Exercise/ or exp counseling/
8.	(rehabilit* or restitut* or remedi* or restorat* or re-train* or retrain* or train* or recover* or treat* or guid* or instruct* or teach* or stimulat* or exerci* or counsel* or therap* or intervent* or manag*).ti,ab.
9.	or/6-8

10.	Cognitive Therapy/ or reminder systems/
11.	neurorehab*.ti,ab.
12.	((perceptual or neuro* or attention or cognitive or memory or cognition or scanning) adj3 (train* or re-train* or retrain* or rehabilit* or interven* or therap*)).ti,ab.
13.	((meta-cognitive or metacognitive or cognitive) adj2 strateg*).ti,ab.
14.	or/10-13
15.	5 and 9
16.	14 or 15

Embase search terms

1.	exp *attention/ or *reinforcement/ or exp *cognition/ or *cognitive defect/ or exp *Memory/ or exp *Memory Disorder/
2.	*neuropsychology/ or *language/ or exp *speech/ or *problem solving/ or *mathematics/ or exp *learning/ or exp *thinking/ or exp *linguistics/
3.	((perceptual or attentional) adj3 (disorders* or deficit* or impairment* or abilit*)).ti,ab.
4.	(cogniti* or neuropsychol* or memor* or learn* or attenti* or information process* or language or visuopat* or visuocognit* or problem solving or reason or execut* or metacognit*).ti,ab.
5.	or/1-4
6.	*Rehabilitation/ or therapy/ or *computer assisted therapy/ or exp *neuropsychological test/
7.	exp *teaching/ or *convalescence/ or exp *Exercise/ or exp *counseling/
8.	(rehabilit* or restitut* or remedi* or restorat* or re-train* or retrain* or train* or recover* or treat* or guid* or instruct* or teach* or stimulat* or exerci* or counsel* or therap* or intervent* or manag*).ti,ab.
9.	or/6-8
10.	5 and 9
11.	*Cognitive Therapy/ or *reminder system/ or *cognitive rehabilitation/
12.	neurorehab*.ti,ab.
13.	((perceptual or neuro* or attention or cognitive or memory or cognition or scanning) adj3 (train* or re-train* or retrain* or rehabilit* or interven* or therap* or rehab*)).ti,ab.
14.	((meta-cognitive or metacognitive or cognitive) adj2 strateg*).ti,ab.
15.	or/10-14

Cochrane search terms

#1.	MeSH descriptor: [Extinction, Psychological] explode all trees
#2.	MeSH descriptor: [Attention] this term only
#3.	MeSH descriptor: [Cognition] explode all trees
#4.	MeSH descriptor: [Cognition Disorders] explode all trees
#5.	MeSH descriptor: [Memory] explode all trees
#6.	MeSH descriptor: [Memory Disorders] explode all trees
#7.	MeSH descriptor: [Neuropsychology] explode all trees
#8.	MeSH descriptor: [Language] explode all trees
#9.	MeSH descriptor: [Speech] explode all trees
#10.	MeSH descriptor: [Problem Solving] explode all trees
#11.	MeSH descriptor: [Mathematics] explode all trees
#12.	MeSH descriptor: [Learning] explode all trees
#13.	MeSH descriptor: [Thinking] explode all trees
#14.	MeSH descriptor: [Psycholinguistics] explode all trees

#15.	((perceptual or attentional) near/3 (disorders* or deficit* or impairment* or abilit*)):ti,ab
#16.	(cogniti* or neuropsychol* or memor* or learn* or attenti* or information process* or language or visuopat* or visuconstruct* or problem solving or reason or execut* or metacognit*):ti,ab
#17.	#1 or #2 or #3 or #4 or #5 or #6 or #7 or #8 or #9 or #10 or #11 or #12 or #13 or #14 or #15 or #16
#18.	MeSH descriptor: [Rehabilitation] explode all trees
#19.	MeSH descriptor: [Therapeutics] explode all trees
#20.	MeSH descriptor: [Neuropsychological Tests] explode all trees
#21.	MeSH descriptor: [Remedial Teaching] explode all trees
#22.	MeSH descriptor: [Therapy, Computer-Assisted] explode all trees
#23.	MeSH descriptor: [Recovery of Function] explode all trees
#24.	MeSH descriptor: [Exercise] explode all trees
#25.	MeSH descriptor: [Counseling] explode all trees
#26.	(rehabilit* or restitut* or remediat* or restorat* or re-train* or retrain* or train* or recover* or treat* or guid* or instruct* or teach* or stimulat* or exerci* or counsel* or therap* or intervent* or manag*):ti,ab
#27.	#18 or #19 or #20 or #21 or #22 or #23 or #24 or #25 or #26
#28.	#17 and #27
#29.	MeSH descriptor: [Cognitive Therapy] explode all trees
#30.	MeSH descriptor: [Reminder Systems] explode all trees
#31.	neurorehab*:ti,ab
#32.	((meta-cognitive or metacognitive or cognitive) next strateg*):ti,ab
#33.	((perceptual or neuro* or attention or cognitive or memory or cognition or scanning) near/3 (train* or re-train* or retrain* or rehabilit* or interven* or therap* or rehab*)):ti,ab
#34.	#28 or #29 or #30 or #31 or #32 or #33

PsycINFO search terms

1.	exp cognition/ or exp cognitive development/ or exp cognitive impairment/ or exp cognitive processes/ or information processing model/ or metacognition/ or need for cognition/
2.	exp comprehension/ or concentration/ or exp concept formation/ or exp decision making/ or naming/ or exp problem solving/ or exp thinking/
3.	neuropsychology/ or exp memory/ or exp memory disorders/ or exp learning/ or exp attention/ or exp visual perception/ or exp language/ or exp mathematical ability/ or exp awareness/
4.	(cogniti* or neuropsychol* or memor* or learn* or attenti* or information process* or language or visuopat* or visuconstruct* or problem solving or reason or execut* or metacognit*):ti,ab.
5.	((perceptual or attentional) adj3 (disorders* or deficit* or impairment* or abilit*)):ti,ab.
6.	or/1-5
7.	exp rehabilitation/ or exp training/ or exp "recovery (disorders)"/ or exp treatment/ or exp cognitive techniques/ or exp intervention/ or exp counseling/ or rehabilitation counseling/
8.	exp teaching/ or exp exercise/ or exp neuropsychological rehabilitation/
9.	(rehabilit* or restitut* or remediat* or restorat* or re-train* or retrain* or train* or recover* or treat* or guid* or instruct* or teach* or stimulat* or exerci* or counsel* or therap* or intervent* or manag* or neurorehab*):ti,ab.
10.	or/7-9
11.	6 and 10
12.	Cognitive Therapy/ or reminder systems/

13.	neurorehab*.ti,ab.
14.	((perceptual or neuro* or attention or cognitive or memory or cognition or scanning) adj3 (train* or re-train* or retrain* or rehabilit* or interven* or {Including Related Terms}))
15.	neurorehab*.ti,ab.
16.	((perceptual or neuro* or attention or cognitive or memory or cognition or scanning) adj3 (train* or re-train* or retrain* or rehabilit* or interven* or therap* or rehab*)).ti,ab.
17.	((meta-cognitive or metacognitive or cognitive) adj2 strateg*).ti,ab.
18.	or/11-17

A.3.14 Non pharmacological treatment

Review for non pharmacological treatment of fatigue, spasticity, mobility, pain ataxia and tremor were run as one search.

For adults with MS, what is the clinical evidence and cost effectiveness of non-pharmacological programmes (including self-management programmes) for:

Fatigue

Spasticity

Mobility

Pain

Ataxia

Tremor

Search constructed by combining the columns in the following table using the AND Boolean operator

Population	Intervention / exposure	Comparison	Study filter used	Date parameters
Adults who have a diagnosis of MS or possible MS, or are being investigated for MS.	As below	n/a	The following filters were used in Medline and Embase only: RCT, SR	See Table 1

Medline search terms

1.	exp Rehabilitation/
2.	exp Physical Therapy Modalities/
3.	Self care/
4.	Self efficacy/
5.	Patient Care Team/
6.	Ambulatory care/
7.	(interdisciplinary or multidisciplinary or inter disciplinary or multi disciplinary or MDT or home based or non pharmacological or non pharma).ti,ab.
8.	((self or own or personal or alone) adj3 (rehab* or efficacy or treatment* or programme* or program* or technique* or manage* or intervention* or therap* or train* or strateg* or method* or counsel* or care* or caring)).ti,ab.
9.	((treatment* or therap* or intervention* or rehab* or fatigue or energy) adj2 (strateg* or method* or programme* or program* or technique* or manage* or train*)).ti,ab.
10.	((energy or fatigue) adj2 (effectiv* or conserv*)).ti,ab.
11.	Transcutaneous Electrical Nerve Stimulation/
12.	(TENS or electroanalgesi* or electro analgesi*).ti,ab.
13.	(electric* Nerve adj2 stimulation adj2 (transcutaneous or percutaneous or analgesi*)).ti,ab.

14.	(electrostimulation adj2 (transcutaneous or percutaneous or analgesi*)).ti,ab.
15.	FACETS.ti,ab.
16.	Mind body therapies/
17.	Relaxation/
18.	Relaxation therapy/
19.	Meditation/
20.	Cognitive Therapy/
21.	(mindfulness or relax* or meditation or cognitive).ti,ab.
22.	Exercise therapies/
23.	Muscle Stretching Exercises/
24.	Physical endurance/
25.	Physical fitness/
26.	exp "Physical Education and Training"/
27.	Exercise Movement Techniques/
28.	(exercising or exercise*).ti,ab.
29.	((physical or muscle* or muscular) adj2 (endurance or fitness or exertion or stretch* or stand* or splinting)).ti,ab.
30.	((exercise or fitness) adj2 (technics or techniques)).ti,ab.
31.	Yoga/ or Tai ji/
32.	(tai ji or tai chi or taichi or taiji or taijiquan).ti,ab.
33.	(gym or calisthenics or pilates or yoga).ti,ab.
34.	Clothing/
35.	lycra.ti,ab.
36.	(cooling adj2 (device* or clothing or clothes or cloth or garment*)).ti,ab.
37.	or/1-36
38.	"Activities of Daily Living"/
39.	(activities or activity or active).ti,ab.
40.	(disability or disable*).ti,ab.
41.	(fatigue* or exhaust* or tired* or weary or weariness or weak* or letharg* or langour or tremor* or ataxia or spasticity or mobil*).ti,ab.
42.	((conserv* or deplet* or low or lack or limited) adj2 energy).ti,ab.
43.	exp Ataxia/
44.	(coordination impairment or ataxy or ataxic or dyscoordination or dyssynergia or incoordination or lack of coordination).ti,ab.
45.	exp Fatigue/
46.	(lassitude or tired or tiring or exhaust*).ti,ab.
47.	exp Muscle weakness/
48.	(weak* adj (muscular or muscle*)).ti,ab.
49.	exp Paraparesis/
50.	muscle spasticity/
51.	(spastic* or spasm*).ti,ab.
52.	Spasm/
53.	Mobility limitation/ or Movement/ or Locomotion/
54.	((limit* or difficult*) adj2 (walk* or ambulation or mobility or moving or movement or locomotion)).ti,ab.
55.	exp Pain/

56.	(pain* or ache or aching or physical suffering).ti,ab.
57.	or/38-56
58.	37 and 57

Embase search terms

1.	exp *rehabilitation/
2.	*self care/
3.	*Patient Care/
4.	*ambulatory care/
5.	(interdisciplinary or multidisciplinary or inter disciplinary or multi disciplinary or MDT or home based or non pharmacological or non pharma).ti,ab.
6.	((self or own or personal or alone) adj3 (rehab* or efficacy or treatment* or programme* or program* or technique* or manage* or intervention* or therap* or train* or strateg* or method* or counsel* or care* or caring)).ti,ab.
7.	((treatment* or therap* or intervention* or rehab* or fatigue or energy) adj2 (strateg* or method* or programme* or program* or technique* or manage* or train*)).ti,ab.
8.	((energy or fatigue) adj2 (effectiv* or conserv*)).ti,ab.
9.	*Transcutaneous Nerve Stimulation/
10.	(TENS or electroanalgesi* or electro analgesi*).ti,ab.
11.	(electric* nerve adj2 stimulation adj2 (transcutaneous or percutaneous or analgesi*)).ti,ab.
12.	(electrostimulation adj2 (transcutaneous or percutaneous or analgesi*)).ti,ab.
13.	FACETS.ti,ab.
14.	*Alternative therapies/
15.	*Physical medicine/
16.	*Relaxation training/
17.	*Meditation/
18.	*Cognitive Therapy/
19.	(mindfulness or relax* or meditation or cognitive).ti,ab.
20.	exp *kinesiotherapy/
21.	*Muscle stretching/
22.	*Stretching Exercises/
23.	*Endurance training/
24.	*Fitness/
25.	exp *Physical Education/
26.	*Movement Therapy/
27.	*Muscle training/
28.	*Exercise/
29.	(exercising or exercise*).ti,ab.
30.	((physical or muscle* or muscular) adj2 (endurance or fitness or exertion or stretch* or stand* or splinting)).ti,ab.
31.	((exercise or fitness) adj2 (technics or techniques)).ti,ab.
32.	*Yoga/ or *Tai ji/
33.	(tai ji or tai chi or taichi or taiji or taijiquan).ti,ab.
34.	(gym or calisthenics or pilates or yoga).ti,ab.
35.	*Clothing/
36.	lycra.ti,ab.
37.	(cooling adj2 (device* or clothing or clothes or cloth or garment*)).ti,ab.

38.	or/1-37
39.	*Daily life activities/
40.	(activities or activity or active).ti,ab.
41.	(disability or disable*).ti,ab.
42.	(fatigue* or exhaust* or tired* or weary or weariness or weak* or letharg* or langour or tremor* or ataxia or ataxic or spasticity or mobil*).ti,ab.
43.	((conserv* or deplet* or low or lack or limited) adj2 energy).ti,ab.
44.	exp *Ataxia/
45.	(coordination impairment or ataxy or dyscoordination or dyssynergia or incoordination or lack of coordination).ti,ab.
46.	exp *fatigue/
47.	(lassitude or tired or tiring or exhaust*).ti,ab.
48.	exp *muscle weakness/
49.	(weak* adj (muscular or muscle*)).ti,ab.
50.	*Paraplegia/
51.	exp *muscle hypertonia/
52.	(spastic* or spasm*).ti,ab.
53.	*muscle spasm/ or *muscle cramp/
54.	*walking difficulty/ or *locomotion/
55.	((limit* or difficult*) adj2 (walk* or ambulation or mobility or moving or movement or locomotion)).ti,ab.
56.	exp *pain/
57.	(pain* or ache or aching or physical suffering).ti,ab.
58.	or/39-57
59.	38 and 58

Cochrane search terms

#1.	MeSH descriptor: [Rehabilitation] explode all trees
#2.	MeSH descriptor: [Physical Therapy Modalities] explode all trees
#3.	MeSH descriptor: [Self Care] explode all trees
#4.	MeSH descriptor: [Self Efficacy] explode all trees
#5.	MeSH descriptor: [Patient Care Team] explode all trees
#6.	MeSH descriptor: [Ambulatory Care] explode all trees
#7.	MeSH descriptor: [Transcutaneous Electric Nerve Stimulation] explode all trees
#8.	MeSH descriptor: [Mind-Body Therapies] explode all trees
#9.	MeSH descriptor: [Relaxation Therapy] explode all trees
#10.	MeSH descriptor: [Relaxation] explode all trees
#11.	MeSH descriptor: [Meditation] explode all trees
#12.	MeSH descriptor: [Cognitive Therapy] explode all trees
#13.	MeSH descriptor: [Exercise Therapy] explode all trees
#14.	MeSH descriptor: [Muscle Stretching Exercises] explode all trees
#15.	MeSH descriptor: [Physical Endurance] explode all trees
#16.	MeSH descriptor: [Physical Fitness] explode all trees
#17.	MeSH descriptor: [Physical Education and Training] explode all trees
#18.	MeSH descriptor: [Exercise Movement Techniques] explode all trees
#19.	MeSH descriptor: [Yoga] explode all trees

#20.	MeSH descriptor: [Tai Ji] explode all trees
#21.	(interdisciplinary or multidisciplinary or inter disciplinary or multi disciplinary or MDT or home based or non pharmacological or non pharma):ti,ab
#22.	((self or own or personal or alone) near/3 (rehab* or efficacy or treatment* or programme* or program* or technique* or manage* or intervention* or therap* or train* or strateg* or method* or counsel* or care* or caring)):ti,ab
#23.	((treatment* or therap* or intervention* or rehab* or fatigue or energy) near/2 (strateg* or method* or programme* or program* or technique* or manage* or train*)):ti,ab
#24.	((energy or fatigue) near/2 (effectiv* or conserv*)):ti,ab
#25.	(TENS or electroanalgesi* or electro analgesi*):ti,ab
#26.	(electric* nerve near/2 stimulation near/2 (transcutaneous or percutaneous or analgesi*)):ti,ab
#27.	(electrostimulation near/2 (transcutaneous or percutaneous or analgesi*)):ti,ab
#28.	FACETS:ti,ab
#29.	(mindfulness or relax* or meditation or cognitive):ti,ab
#30.	((physical or muscle* or muscular) near/2 (endurance or fitness or exertion or stretch* or stand* or splinting)):ti,ab
#31.	((exercise or fitness) near/2 (technics or techniques)):ti,ab
#32.	(tai ji or tai chi or taichi or taiji or taijiquan):ti,ab
#33.	(gym or calisthenics or pilates or yoga):ti,ab
#34.	lycra:ti,ab
#35.	(cooling near/2 (device* or clothing or clothes or cloth or garment*)):ti,ab
#36.	{or #1-#35}
#37.	MeSH descriptor: [Activities of Daily Living] explode all trees
#38.	MeSH descriptor: [Ataxia] explode all trees
#39.	MeSH descriptor: [Fatigue] explode all trees
#40.	MeSH descriptor: [Muscle Weakness] explode all trees
#41.	MeSH descriptor: [Paraparesis] explode all trees
#42.	MeSH descriptor: [Muscle Spasticity] explode all trees
#43.	MeSH descriptor: [Spasm] explode all trees
#44.	MeSH descriptor: [Mobility Limitation] explode all trees
#45.	MeSH descriptor: [Movement] explode all trees
#46.	MeSH descriptor: [Locomotion] explode all trees
#47.	MeSH descriptor: [Pain] explode all trees
#48.	(activities or activity or active):ti,ab
#49.	(disability or disable*):ti,ab
#50.	(fatigue* or exhaust* or tired* or weary or weariness or weak* or letharg* or langour or tremor* or ataxia or spasticity or mobil*):ti,ab
#51.	((conserv* or deplet* or low or lack or limited) near/2 energy):ti,ab
#52.	(coordination impairment or ataxy or ataxic or dyscoordination or dyssynergia or incoordination or lack of coordination):ti,ab
#53.	(lassitude or tired or tiring or exhaust*):ti,ab
#54.	(weak* near (muscular or muscle*)):ti,ab
#55.	(spastic* or spasm*):ti,ab
#56.	((limit* or difficult*) near/2 (walk* or ambulation or mobility or moving or movement or locomotion)):ti,ab
#57.	(pain* or ache or aching or physical suffering):ti,ab

#58.	{or #37-#57}
#59.	#36 and #57

A.3.15 Rehabilitation settings

For adults with MS, what is the clinical evidence and cost effectiveness of rehabilitation provided in different settings?

Search constructed by combining the columns in the following table using the AND Boolean operator

Population	Intervention / exposure	Comparison	Study filter used	Date parameters
Adults who have a diagnosis of MS or possible MS, or are being investigated for MS. Including CIS.	As below	n/a	The following filters were used in Medline and Embase only: RCT, SR, OBS	See Table 1

Medline search terms

1.	exp Multiple Sclerosis/rh [Rehabilitation]
2.	exp Rehabilitation/
3.	(neurorehab\$ or rehabilitat\$ or rehab).ti,ab.
4.	"Recovery of Function"/
5.	Rehabilitation Centers/
6.	Rehabilitation Nursing/
7.	or/1-6
8.	exp Physical Therapy Modalities/
9.	(physiotherap\$ or physical therap\$ or range of motion or exercise\$ or physical fitness).ti,ab.
10.	((strength\$ or muscle\$ or stretch\$ or resistance or balance or mobility or resistive or gait) adj2 training).ti,ab.
11.	"Physical Education and Training"/
12.	Physical Fitness/
13.	or/8-12
14.	(occupational therap\$ or assistive devices).ti,ab.
15.	((personal or day or daily) adj3 activit\$).ti,ab.
16.	Self Care/
17.	((pacing or pace or mobility or coping or bathing or dressing or grooming or washing or toilet\$ or eating) adj2 (activit\$ or technique\$ or skill\$ or assist\$ or aid\$ or train\$)).ti,ab.
18.	(wellness adj2 (program\$ or intervention\$ or training)).ti,ab.
19.	or/14-18
20.	Language Disorders/rh [Rehabilitation]
21.	((speech or language or swallow\$ or aphasia or dysphagia) adj2 (therap\$ or train\$ or remed\$)).ti,ab.
22.	or/20-21
23.	Employment, Supported/
24.	((occupation\$ or employ\$ or vocation\$ or job\$ or work or workplace) adj2 (return\$ or retrain\$ or support\$ or readiness or coaching or rehabilit\$)).ti,ab.
25.	or/23-24
26.	psychotherapy/ or cognitive therapy/

27.	Behavior Therapy/
28.	((cbt or (cognitive or psycholog\$ or psychosocial or behavio?r\$)) adj2 (therap\$ or support or intervention\$ or treatment\$)).ti,ab.
29.	((memory or remind\$ or remember\$) adj2 (training or retrain\$ or rehab\$ or skill\$)).ti,ab.
30.	psychotherap\$.ti,ab.
31.	Memory Disorders/rh [Rehabilitation]
32.	or/26-31
33.	7 or 13 or 19 or 22 or 25 or 32
34.	*patient care/ or ambulatory care/
35.	inpatients/ or outpatients/
36.	Patient Care Team/
37.	Hospitalization/
38.	community health services/ or home care services/ or home care services, hospital-based/ or Rehabilitation Centers/
39.	"Delivery of Health Care"/
40.	hospital units/ or self-care units/
41.	(outpatient\$ or inpatient\$).ti,ab.
42.	((community or home or hospital) adj2 (based or setting\$ or care\$ or service\$ or unit\$)).ti,ab.
43.	((multi?disciplinary or inter?disciplinary or cross?disciplinary or integrated or multi?modal or multi?professional or inter?professional or cross?professional) adj3 (team\$ or rehab\$ or care\$ or therap\$)).ti,ab.
44.	(MDT and (rehab\$ or neurorehab\$)).ti,ab.
45.	(rehab\$ adj4 setting\$).ti,ab.
46.	(residential adj2 (care\$ or setting\$ or unit\$)).ti,ab.
47.	((family or families or carer\$ or care?giver\$) adj3 (setting\$ or care or center\$ or centre\$ or support\$ or therap\$ or based or centric or focus\$ or intervention\$)).ti,ab.
48.	((group\$ or individual\$) adj3 (intervention\$ or session\$ or workshop\$ or rehab\$ or program\$ or therap\$)).ti,ab.
49.	or/34-48
50.	33 and 49

Embase search terms

1.	multiple sclerosis/rh [Rehabilitation]
2.	*rehabilitation/ or cognitive rehabilitation/ or community based rehabilitation/ or functional assessment/ or functional training/ or home rehabilitation/ or muscle training/ or occupational therapy/ or psychosocial rehabilitation/ or recreational therapy/ or exp speech rehabilitation/ or vocational rehabilitation/
3.	rehabilitation care/ or rehabilitation center/ or rehabilitation nursing/ or rehabilitation patient/
4.	(neurorehab\$ or rehabilitat\$ or rehab).ti,ab.
5.	or/1-4
6.	physiotherapy/ or home physiotherapy/
7.	fitness/
8.	physical education/
9.	(physiotherap\$ or physical therap\$ or range of motion or exercise\$ or physical fitness).ti,ab.
10.	((strength\$ or muscle\$ or stretch\$ or resistance or balance or mobility or resisitive or gait) adj2 training).ti,ab.

11.	or/6-10
12.	(occupational therap\$ or assistive devices).ti,ab.
13.	((personal or day or daily) adj3 activit\$).ti,ab.
14.	self care/ or self help/
15.	((pacing or pace or coping or mobility or bathing or dressing or grooming or washing or toilet\$ or eating) adj2 (activit\$ or technique\$ or skill\$ or assist\$ or aid\$ or train\$)).ti,ab.
16.	(wellness adj2 (program\$ or intervention\$ or training)).ti,ab.
17.	daily life activity/
18.	or/12-17
19.	exp speech disorder/rh [Rehabilitation]
20.	speech therapy/
21.	((speech or language or swallow\$ or aphasia or dysphagia) adj2 (therap\$ or train\$ or remed\$)).ti,ab.
22.	or/19-21
23.	exp employment/
24.	((occupation\$ or employ\$ or vocation\$ or job\$ or work or workplace) adj2 (return\$ or retrain\$ or support\$ or readiness or coaching or rehabilit\$)).ti,ab.
25.	23 or 24
26.	psychotherapy/ or behavior therapy/ or cognitive therapy/
27.	((cbt or (cognitive or psycholog\$ or psychosocial or behavio?r\$)) adj2 (therap\$ or support or intervention\$ or treatment\$)).ti,ab.
28.	((memory or remind\$ or remember\$) adj2 (training or retrain\$ or rehab\$ or skill\$)).ti,ab.
29.	psychotherap\$.ti,ab.
30.	exp memory disorder/rh [Rehabilitation]
31.	or/26-30
32.	5 or 11 or 18 or 22 or 25 or 31
33.	community based rehabilitation/ or home rehabilitation/
34.	home physiotherapy/
35.	hospital patient/
36.	outpatient/ or outpatient care/
37.	hospitalization/
38.	exp ambulatory care/
39.	*patient care/ or rehabilitation care/
40.	rehabilitation center/
41.	community health nursing/
42.	community care/
43.	home care/
44.	*health care delivery/
45.	"hospital subdivisions and components"/
46.	(outpatient\$ or inpatient\$).ti,ab.
47.	((community or home or hospital) adj2 (based or setting\$ or care\$ or service\$ or unit\$)).ti,ab.
48.	((multi?disciplinary or inter?disciplinary or cross?disciplinary or integrated or multi?modal or multi?professional or inter?professional or cross?professional) adj3 (team\$ or rehab\$ or care\$ or therap\$)).ti,ab.
49.	(MDT and (rehab\$ or neurorehab\$)).ti,ab.
50.	(rehab\$ adj4 setting\$).ti,ab.

51.	(residential adj2 (care\$ or setting\$ or unit\$)).ti,ab.
52.	((family or families or carer\$ or care?giver\$) adj3 (setting\$ or care or center\$ or centre\$ or support\$ or therap\$ or based or centric or focus\$ or intervention\$)).ti,ab.
53.	((group\$ or individual\$) adj3 (intervention\$ or session\$ or workshop\$ or rehab\$ or program\$ or therap\$)).ti,ab.
54.	or/33-53
55.	32 and 54

Cochrane search terms

#1.	MeSH descriptor: [Rehabilitation] explode all trees
#2.	MeSH descriptor: [Multiple Sclerosis] explode all trees and with qualifier(s): [Rehabilitation - RH]
#3.	(neurorehab* or rehabilitat* or rehab):ti,ab
#4.	MeSH descriptor: [Recovery of Function] this term only
#5.	MeSH descriptor: [Rehabilitation Centers] this term only
#6.	MeSH descriptor: [Rehabilitation Nursing] this term only
#7.	MeSH descriptor: [Physical Therapy Modalities] explode all trees
#8.	MeSH descriptor: [Physical Education and Training] this term only
#9.	MeSH descriptor: [Physical Fitness] this term only
#10.	(physiotherap* or exercise* or "physical therapy" or "physical therapies" or "range of motion" or "physical fitness"):ti,ab
#11.	((strength* or muscle* or stretch* or resistance or resistive or balance or mobility or gait) near/2 training):ti,ab
#12.	(occupation* next/1 therap*):ti,ab
#13.	assistive devices:ti,ab
#14.	((personal or day or daily) near/3 activit*):ti,ab
#15.	MeSH descriptor: [Self Care] this term only
#16.	((pacing or pace or mobility or coping or bathing or dressing or grooming or washing or toilet* or eating) near/2 (activit* or technique* or skill* or assist* or aid* or train*)):ti,ab
#17.	(wellness near/2 (program* or intervention* or training*)):ti,ab
#18.	MeSH descriptor: [Speech Disorders] explode all trees and with qualifier(s): [Rehabilitation - RH]
#19.	((speech or language or swallow* or aphasia or dysphagia) near/2 (therap* or train* or remed*)):ti,ab
#20.	MeSH descriptor: [Employment, Supported] this term only
#21.	((occupation* or employ* or vocation* or job* or work* or workplace) near/2 (return* or retrain* or support* or readiness or coaching or rehabilit*)):ti,ab
#22.	MeSH descriptor: [Psychotherapy] this term only
#23.	MeSH descriptor: [Cognitive Therapy] this term only
#24.	MeSH descriptor: [Behavior Therapy] this term only
#25.	CBT:ti,ab
#26.	((cognitive or psycholog* or psychosocial* or behavior* or behaviour*) near/2 (therap* or support or intervention* or treatment*)):ti,ab
#27.	((memory or remind* or remember*) near/2 (training or retrain* or rehab* or skill*)):ti,ab
#28.	psychotherap*:ti,ab
#29.	MeSH descriptor: [Memory Disorders] this term only and with qualifier(s): [Rehabilitation - RH]
#30.	#1 or #2 or #3 or #4 or #5 or #6 or #7 or #8 or #9 or #10 or #11 or #12 or #13 or #14 or #15 or #16 or #17 or #18 or #19 or #20 or #21 or #22 or #23 or #24 or #25 or #26 or #27 or #28 or #29

#31.	MeSH descriptor: [Patient Care] this term only
#32.	MeSH descriptor: [Ambulatory Care] this term only
#33.	MeSH descriptor: [Inpatients] this term only
#34.	MeSH descriptor: [Outpatients] this term only
#35.	MeSH descriptor: [Patient Care Team] this term only
#36.	MeSH descriptor: [Hospitalization] this term only
#37.	MeSH descriptor: [Community Health Services] this term only
#38.	MeSH descriptor: [Home Care Services] this term only
#39.	MeSH descriptor: [Home Care Services, Hospital-Based] this term only
#40.	MeSH descriptor: [Rehabilitation Centers] this term only
#41.	MeSH descriptor: [Delivery of Health Care] this term only
#42.	MeSH descriptor: [Hospital Units] this term only
#43.	MeSH descriptor: [Self-Care Units] this term only
#44.	(inpatient* or outpatient*):ti,ab
#45.	((community or home or hospital) near/2 (based or setting* or care* or service* or unit*)):ti,ab
#46.	((multi?disciplinary or inter?disciplinary or cross?disciplinary or integrated or multi?modal or multi?professional or inter?professional or cross?professional) near/2 (team* or rehab* or care* or therap*)):ti,ab
#47.	(MDT and (rehab* or neurorehab*)):ti,ab
#48.	(rehab* near/4 setting*):ti,ab
#49.	(residential near/2 (care* or setting* or unit*)):ti,ab
#50.	((family or families or carer* or care?giver*) near/3 (setting* or care* or center* or centre* or support* or therap* or based or centric or focus* or intervention*)):ti,ab
#51.	((group* or individual*) near/3 (intervention* or session* or workshop* or rehab* or program* or therap*)):ti,ab
#52.	#31 or #32 or #33 or #34 or #35 or #36 or #37 or #38 or #39 or #40 or #41 or #42 or #43 or #44 or #45 or #46 or #47 or #48 or #49 or #50 or #51
#53.	#30 and #52

PsycINFO search terms

1.	rehabilitation/ or cognitive rehabilitation/ or exp neuropsychological rehabilitation/ or neurorehabilitation/ or occupational therapy/ or physical therapy/ or exp psychosocial rehabilitation/ or "activities of daily living"/ or adaptive behavior/ or disability management/ or rehabilitation centers/ or self care skills/
2.	(neurorehab\$ or rehabilitat\$ or rehab).ti,ab.
3.	physical education/ or physical fitness/ or physical mobility/
4.	(physiotherap\$ or physical therap\$ or range of motion or exercise\$ or physical fitness).ti,ab.
5.	((strength\$ or muscle\$ or stretch\$ or resistance or balance or mobility or resisitive or gait) adj2 training).ti,ab.
6.	(occupational therap\$ or assistive devices).ti,ab.
7.	((personal or day or daily) adj3 activit\$).ti,ab.
8.	((pacing or pace or coping or bathing or dressing or grooming or washing or toilet\$ or eating or mobility) adj2 (activit\$ or technique\$ or assist\$ or aid\$ or train\$)).ti,ab.
9.	(wellness adj2 (program\$ or intervention\$ or training)).ti,ab.
10.	Speech Therapy/
11.	((speech or language or aphasia or dysphagia) adj2 (therap\$ or train\$ or remed\$)).ti,ab.
12.	supported employment/ or vocational rehabilitation/ or employability/

13.	((occupation\$ or employ\$ or vocation\$ or job\$ or work or workplace) adj2 (return\$ or retrain\$ or support\$ or readiness or coaching or rehabilit\$)).ti,ab.
14.	psychotherapy/ or exp behavior therapy/ or exp cognitive behavior therapy/ or cognitive therapy/
15.	memory training/
16.	((cbt or (cognitive or psycholog\$ or psychosocial or behavio?r\$)) adj2 (therap\$ or support or intervention\$ or treatment\$)).ti,ab.
17.	((memory or remind\$ or remember\$) adj2 (training or retrain\$ or rehab\$ or skill\$)).ti,ab.
18.	psychotherap\$.ti,ab.
19.	or/1-18
20.	health care services/
21.	outpatient treatment/ or home care/ or outpatients/
22.	hospitalization/
23.	Hospitalized Patients/
24.	Health Care Delivery/
25.	exp Interdisciplinary Treatment Approach/
26.	community services/
27.	rehabilitation centers/
28.	hospital programs/
29.	(outpatient\$ or inpatient\$).ti,ab.
30.	((community or home or hospital) adj2 (based or setting\$ or care\$ or service\$ or unit\$)).ti,ab.
31.	((multi?disciplinary or inter?disciplinary or cross?disciplinary or integrated or multi?modal or multi?professional or inter?professional or cross?professional) adj3 (team\$ or rehab\$ or care\$ or therap\$)).ti,ab.
32.	(MDT and (rehab\$ or neurorehab\$)).ti,ab.
33.	(rehab\$ adj4 setting\$).ti,ab.
34.	(residential adj2 (care\$ or setting\$ or unit\$)).ti,ab.
35.	((family or families or carer\$ or care?giver\$) adj3 (setting\$ or care or center\$ or centre\$ or support\$ or therap\$ or based or centric or focus\$ or intervention\$)).ti,ab.
36.	((group\$ or individual\$) adj3 (intervention\$ or session\$ or workshop\$ or rehab\$ or program\$ or therap\$)).ti,ab.
37.	group intervention/
38.	or/20-37
39.	19 and 38

A.3.16 Vitamin D

For adults with MS, what is the clinical evidence and cost effectiveness of pharmacological management with vitamin D?

Search constructed by combining the columns in the following table using the AND Boolean operator

Population	Intervention / exposure	Comparison	Study filter used	Date parameters
Adults who have a diagnosis of MS or possible MS, or are being investigated for MS.	As below	n/a	The following filters were used in Medline and Embase only: RCT, SR	See Table 1

Medline search terms

1.	exp Vitamin D/
2.	(vitamin adj (D or D2 or D3 or D4 or D5)).ti,ab.
3.	(paracalcitol* or Zemplar* or ergocalciferol* or alfacalcidol* or One-Alpha* or calcitriol* or Rocaltrol* or Calcijex* or oxacalcitriol* or falecalcitriol* or fluorocalcitril*).ti,ab.
4.	(dihydrotachysterol* or maxacalcitol* or calciferol* or calcifediol* or doxercalciferol* or cholecalciferol* or ercalcidiol* or hectorol* or sitocalciferol* or paracalcin*).ti,ab.
5.	(dihydroxyvitamin* or hydroxyvitamin* or hydroxycalciferol* or dihydroxycalciferol* or hydroxyergocalciferol* or dihydroxyergocalciferol* or hydroxycholecalciferol* or dihydroxycholecalciferol*).ti,ab.
6.	or/1-5

Embase search terms

1.	exp Vitamin D/
2.	(vitamin adj (D or D2 or D3 or D4 or D5)).ti,ab.
3.	(paracalcitol* or Zemplar* or ergocalciferol* or alfacalcidol* or One-Alpha* or calcitriol* or Rocaltrol* or Calcijex* or oxacalcitriol* or falecalcitriol* or fluorocalcitril*).ti,ab.
4.	(dihydrotachysterol* or maxacalcitol* or calciferol* or calcifediol* or doxercalciferol* or cholecalciferol* or ercalcidiol* or hectorol* or sitocalciferol* or paracalcin*).ti,ab.
5.	(dihydroxyvitamin* or hydroxyvitamin* or hydroxycalciferol* or dihydroxycalciferol* or hydroxyergocalciferol* or dihydroxyergocalciferol* or hydroxycholecalciferol* or dihydroxycholecalciferol*).ti,ab.
6.	or/1-5

Cochrane search terms

#1.	MeSH descriptor: [Vitamin D] explode all trees
#2.	(vitamin next (D or D2 or D3 or D4 or D5)):ti,ab
#3.	(paracalcitol* or Zemplar* or ergocalciferol* or alfacalcidol* or One-Alpha* or calcitriol* or Rocaltrol* or Calcijex* or oxacalcitriol* or falecalcitriol* or fluorocalcitril*).ti,ab
#4.	(dihydrotachysterol* or maxacalcitol* or calciferol* or calcifediol* or doxercalciferol* or cholecalciferol* or ercalcidiol* or hectorol* or sitocalciferol* or paracalcin*).ti,ab
#5.	(dihydroxyvitamin* or hydroxyvitamin* or hydroxycalciferol* or dihydroxycalciferol* or hydroxyergocalciferol* or dihydroxyergocalciferol* or hydroxycholecalciferol* or dihydroxycholecalciferol*).ti,ab
#6.	#1 or #2 or #3 or #4 or #5

A.3.17 Complementary therapies

Review for omegas and acupuncture were run as one search

For adults with MS, what is the clinical evidence and cost effectiveness of non-pharmacological management with complementary and alternative therapies (omega-3 fatty acids, omega-6 fatty acids, acupuncture)?

Search constructed by combining the columns in the following table using the AND Boolean operator

Population	Intervention / exposure	Comparison	Study filter used	Date parameters
Adults who have a diagnosis of MS or possible MS, or are being investigated for MS.	As below	n/a	The following filters were used in Medline and Embase only: RCT, SR, OBS	See Table 1

Medline search terms

1.	Acupuncture/ or acupressure/
2.	exp Acupuncture Therapy/
3.	Electroacupuncture/
4.	(acupunctur\$ or electroacupunctur\$ or acupressur\$).ti,ab.
5.	exp Fish Oils/
6.	fatty acids, unsaturated/ or fatty acids, essential/ or exp fatty acids, omega-3/ or exp fatty acids, omega-6/
7.	exp Dietary Fats, Unsaturated/
8.	((fish or soybean or soy bean or soya or palm or rapeseed or primrose or linseed or flax or cod liver or safflower or sunflower or corn or maize or sesame or cottonseed or cotton seed) adj3 oil\$).ti,ab.
9.	(omega 3 or omega-3 or omega-6 or omega 6).ti,ab.
10.	PUFA\$.ti,ab.
11.	((n3 or n-3 or essential or unsaturated or polyunsaturated or n6 or n-6) adj3 (fatty adj3 acid\$)).ti,ab.
12.	((docosahex?enoic or eicosapent?enoic or icosapent?enoic or timnodonic or linolenic or linoleic) adj2 acid\$).ti,ab.
13.	(flaxseed\$ or flax seed\$).ti,ab.
14.	((DHA or ALA or EPA) and (omega-3 or omega-6 or omega 3 or omega 6 or PUFA\$ or fatty acid\$)).ti,ab.
15.	or/1-14

Embase search terms

1.	Acupuncture/ or acupressure/
2.	exp Acupuncture Therapy/
3.	Electroacupuncture/
4.	(acupunctur\$ or electroacupunctur\$ or acupressur\$).ti,ab.
5.	exp Fish Oils/
6.	fatty acids, unsaturated/ or fatty acids, essential/ or exp fatty acids, omega-3/ or exp fatty acids, omega-6/
7.	exp Dietary Fats, Unsaturated/
8.	((fish or soybean or soy bean or soya or palm or rapeseed or primrose or linseed or flax or cod liver or safflower or sunflower or corn or maize or sesame or cottonseed or cotton seed) adj3 oil\$).ti,ab.
9.	(omega 3 or omega-3 or omega-6 or omega 6).ti,ab.
10.	PUFA\$.ti,ab.
11.	((n3 or n-3 or essential or unsaturated or polyunsaturated or n6 or n-6) adj3 (fatty adj3 acid\$)).ti,ab.
12.	((docosahex?enoic or eicosapent?enoic or icosapent?enoic or timnodonic or linolenic or linoleic) adj2 acid\$).ti,ab.
13.	(flaxseed\$ or flax seed\$).ti,ab.
14.	((DHA or ALA or EPA) and (omega-3 or omega-6 or omega 3 or omega 6 or PUFA\$ or fatty acid\$)).ti,ab.
15.	1-14

Cochrane search terms

#1.	MeSH descriptor: [Acupuncture] this term only
#2.	MeSH descriptor: [Acupuncture Therapy] explode all trees

#3.	(acupunctur* or electroacupunctur* or acupressur*):ti,ab
#4.	MeSH descriptor: [Acupressure] this term only
#5.	MeSH descriptor: [Fish Oils] explode all trees
#6.	MeSH descriptor: [Fatty Acids, Unsaturated] this term only
#7.	MeSH descriptor: [Fatty Acids, Essential] this term only
#8.	MeSH descriptor: [Fatty Acids, Omega-3] explode all trees
#9.	MeSH descriptor: [Fatty Acids, Omega-6] explode all trees
#10.	MeSH descriptor: [Linolenic Acids] explode all trees
#11.	MeSH descriptor: [Dietary Fats, Unsaturated] explode all trees
#12.	((fish or soybean or "soy bean" or soya or palm or rapeseed or primrose or linseed or flax or "cod liver" or safflower or sunflower or corn or maize or sesame or cottonseed or "cotton seed") next/2 oil*):ti,ab
#13.	(omega-3 or "omega 3" or "omega 6" or omega-6):ti,ab
#14.	PUFA*:ti,ab
#15.	((n3 or N-3 or "N 3" or essential or unsaturated or polyunsaturated or n6 or n-6 or "n 6") next/2 ("fatty acid" or "fatty acids")):ti,ab
#16.	(flaxseed* or "flax seed" or "flax seeds"):ti,ab
#17.	((docosahex?enoic or eicosapent?enoic or icosapent?enoic or timnodonic or linolenic or linoleic) next/2 acid*):ti,ab
#18.	((DHA or ALA or EPA) and (omega-3 or omega-6 or "omega 3" or "omega 6" or PUFA* or "fatty acid" or "fatty acids")):ti,ab
#19.	#1 or #2 or #3 or #4 or #5 or #6 or #7 or #8 or #9 or #10 or #11 or #12 or #13 or #14 or #15 or #16 or #17 or #18

Amed search terms

1.	acupuncture/
2.	exp acupuncture therapy/
3.	(acupunctur\$ or electroacupunctur\$ or acupressur\$).ti,ab.
4.	fish oils/
5.	fatty acids/
6.	(flaxseed\$ or flax seed\$).ti,ab.
7.	((DHA or ALA or EPA) and (omega-3 or omega-6 or omega 3 or omega 6 or PUFA\$ or fatty acid\$)).ti,ab.
8.	((docosahex?enoic or eicosapent?enoic or icosapent?enoic or timnodonic or linolenic or linoleic) adj2 acid\$).ti,ab.
9.	((fish or soybean or soy bean or soya or palm or rapeseed or primrose or linseed or flax or cod liver or safflower or sunflower or corn or maize or sesame or cottonseed or cotton seed) adj3 oil\$).ti,ab.
10.	(omega 3 or omega-3 or omega-6 or omega 6).ti,ab.
11.	PUFA\$.ti,ab.
12.	((n3 or n-3 or essential or unsaturated or polyunsaturated or n6 or n-6) adj3 (fatty adj3 acid\$)).ti,ab.
13.	or/1-12

A.4 Health economics searches

A.4.1 Economic reviews

Economic searches were conducted in Medline, Embase, HEED and CRD for NHS EED and HTA.

Population	Intervention / exposure	Comparison	Study filter used	Date parameters
Adults who have a diagnosis of MS or possible MS, or are being investigated for MS.	n/a	n/a	The following filters were used in Medline and Embase only: HE	Medline and Embase 2009-13/2/14 HEED, CRD EED and HTA No restriction, search run to 13/2/14

CRD search terms

#1.	MeSH DESCRIPTOR multiple sclerosis EXPLODE ALL TREES WITH QUALIFIER undefined
#2.	(multiple):TI OR (disseminated):TI IN NHSEED, HTA
#2.	(sclero*):TI IN NHSEED, HTA
#3.	#2 AND #3
#4.	#1 OR #4

HEED search terms

1.	ax=multiple and scleros*
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A.4.2 Quality of life reviews

Quality of life searches were conducted in Medline and Embase.

Population	Intervention / exposure	Comparison	Study filter used	Date parameters
Adults who have a diagnosis of MS or possible MS, or are being investigated for MS.	n/a	n/a	The following filters were used in Medline and Embase only: QOL	See table 1

A.4.3 Additional economic search

An additional economics search for Baclofen, section A.3.12, was conducted in Medline, Embase, HEED and CRD for NHS EED and HTA.

Population	Intervention / exposure	Comparison	Study filter used	Date parameters
No population	As below	n/a	The following filters were used in Medline and Embase only: HE	Medline and Embase 2009-13/2/14 HEED, CRD EED and HTA No restriction, search run to 13/2/14

CRD search terms

#1.	MeSH DESCRIPTOR baclofen EXPLODE ALL TREES IN NHSEED,HTA
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#2.	(baclofen):TI OR (baclofen):SO IN NHSEED, HTA
#3.	(lioresal):TI OR (lioresal):SO IN NHSEED, HTA
#4	#1 OR #2 OR #3

HEED search terms

1.	ax=baclofen
2.	ax=lioresal
3.	cs=1 or 2

Medline search terms

1.	Baclofen/
2.	(baclofen* or lioresal).ti,ab.
3.	1 or 2

Embase search terms

1.	Baclofen/
2.	baclofen*.ti,ab.
#3.	1 or 2