National Institute for Health and Care Excellence

Final

Epilepsies in children, young people and adults

[A] Magnetic resonance imaging scan to detect relevant abnormalities in people with epilepsy

NICE guideline NG217

Evidence reviews underpinning recommendations 1.3.1 to 1.3.7 in the NICE guideline

April 2022

Final

These evidence reviews were developed by the National Guideline Alliance which is a part of the Royal College of Obstetricians and Gynaecologists



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Evidence review for magnetic resonance imaging scan to detect relevant abnormalities in people with epilepsy

Review question

What is the yield of relevant abnormalities detected by MRI in people with epilepsy?

Introduction

Magnetic resonance imaging (MRI) enables detailed images based on the effect of magnetic fields on water molecules in the brain. It enables very detailed pictures to be obtained, and utilising different sequences we can gain information about structural abnormalities that could be a cause of epilepsy. Sequences are optimised to enable maximal contrast between grey and white matter, to obtain accurate pictures of the cerebral cortex, the likely area from where epileptic seizures arise. It is the imaging technique of choice in the investigation of people with epilepsy. The aim of this review is to assess how well MRI performns in detecting brain lesions or other relevant abnormalities in people with epilepsy. Knowing the proportion of epilepsy related (clinically relevant abnormalities) and non-epielpsy related abnormalities detected by MRI helps clinicians to recognise those people who are most at risk of adverse outcomes. Information from MRI is used to optimise therapeutic options, and may help to determine who would benefit of surgery for controlling seizures.

Summary of the protocol

Please see Table 1 for a summary of the Population, Intervention, Comparison and Outcome (PICO) characteristics of this review.

Table 1: Summary of the protocol (PICO table)

Population	People with 1 or more confirmed epileptic seizures
Intervention	Magnetic resonance imaging (MRI)
Comparison	Not relevant
Outcomes	Primary outcomes • Proportion identified with a clinically relevant abnormality Secondary outcomes • Proportion identified with a non-epilepsy related abnormality

For further details see the review protocol in appendix A.

Methods and process

This evidence review was developed using the methods and process described in <u>Developing NICE guidelines: the manual</u>. Methods specific to this review question are described in the review protocol in appendix A and the methods document (supplementary document 1).

Declarations of interest were recorded according to NICE's conflicts of interest policy.

Clinical evidence

Included studies

Thirty-nine observational studies (prospective/retrospective single-arm, cohort and cross-sectional studies) were identified for inclusion in this review (Alam-Eldeen 2015, Ali 2017, Asadi-Pooya 2012, Aslan 2010, Bakhsh 2013, Benson 2019, Berg 2000, Betting 2006, Bruno 2017, Byars 2007, Coryell 2018, Craven 2012, Das 2013, Dirik 2018, Dura-Trave 2012, Ekici 2013, Ferreira 2004, Gaillard 2007, Griffiths 2005, Hakami 2013, Harini 2018, Hesdorffer 2008, Hnojcikova 2010, Hsieh 2010, Jasim 2018, Jeniffer 2015, Koirala 2011, Labate 2006, Lefkopoulos 2005, Ma 2019, Nair 2009, Petrou 2007, Rasool 2012, Santos 2005, Sinha 2012, Solosrungruang 2007, Toledo 2013, Wieshmann 2003, Wongladarom 2004).

MRI abnormalities were categorised into various groups including congenital/developmental abnormalities, tumours and vascular pathology (see appendix M for full list). Although exact causality could not be established from the studies, these abnormalities were divided into 'epilepsy related' (this is, clinically relevant hereafter) and 'non-epilepsy related' based on whether or not the lesions were likely to be associated with or cause epilepsy. Examples of clinically relevant abnormalities include malformations of cortical development, tumours, vascular malformations, metabolic/genetic syndromes and acquired lesions such as infection. Examples of non-epilepsy related abnormalities include arachnoid cysts and hydrocephalus which, although there are rare reports of them causing epilepsy, are for the large part incidental findings.

Analyses were not split by MRI type/technology because no studies were identified reporting data on both MRI and CT, however a separate evidence report was produced assessing the yield of relevant abnormalities detected by CT scans in people with epilepsy (see evidence report B).

The included studies are summarised in Table 2.

See the literature search strategy in appendix B and study selection flow chart in appendix C.

Excluded studies

Studies not included in this review with reasons for their exclusions are provided in appendix K.

Summary of clinical studies included in the evidence review

Summaries of the studies that were included in this review are presented in Table 2.

Table 2: Summary of included studies

Study	Population	Intervention	Outcomes
Alam-Eldeen 2015 Retrospective cohort study Egypt	N=89 children with epilepsy from the general population Age at follow up, years, mean (range): 4.3 (1 month to 17 years)	• MRI 1.5-t	 Proportion identified with a clinically relevant abnormality Proportion identified with a non-epilepsy related abnormality

Study	Population	Intervention	Outcomes
Ali 2017 Cross-sectional Pakistan	N=209 people with epilepsy from the general population No demographic characteristics were reported	• MRI 1.5-t	 Proportion identified with a clinically relevant abnormality Proportion identified with a non-epilepsy related abnormality
Asadi-Pooya 2012 Cross-sectional Iran	N=135 children with Lennox- Gastaut syndrome Age at follow-up, years, mean (SD): 3.2 (3.8)	• MRI 1.5-t	 Proportion identified with a clinically relevant abnormality Proportion identified with a non-epilepsy related abnormality
Aslan 2010 Retrospective cohort Turkey	N=32 young people with genetic (idiopathic) generalised epilepsy Age at follow-up, years, mean (range): 22 (16 to 37)	• MRI 1.5-t	 Proportion identified with a clinically relevant abnormality Proportion identified with a non-epilepsy related abnormality
Bakhsh 2013 Prospective cohort Pakistan	N=44 young people with genetic (idiopathic) generalised epilepsy Age at follow-up, years, mean (SD): 19.5 (SD not reported)	• MRI 1-t	 Proportion identified with a clinically relevant abnormality Proportion identified with a non-epilepsy related abnormality
Benson 2019 Retrospective cohort US	N=57 adults with unruptured intracranial arteriovenous malformations associated with seizures [∆] Age at follow-up, years, mean (SD): 35.9 (SD not reported)	• MRI 1.5 or 3.0-t	 Proportion identified with a clinically relevant abnormality Proportion identified with a non-epilepsy related abnormality
Berg 2000 Retrospective cohort US	N=388 children with newly diagnosed epilepsy Age at seizure onset, years, median (IQR): 5.7 (IQR not reported)	MRI (strength of magnet not reported)	 Proportion identified with a clinically relevant abnormality Proportion identified with a non-epilepsy related abnormality
Betting 2006 Prospective cohort Brazil	N=134 adults with genetic (idiopathic) generalised epilepsy Age at seizure onset, years, mean (SD): 28 (9)	• MRI 2-t	 Proportion identified with a clinically relevant abnormality Proportion identified with a non-epilepsy related abnormality

Study	Population	Intervention	Outcomes
	Age at follow up, years, mean (SD): 13 (7)		
Bruno 2017 Prospective cohort Bhutan	N=217 people with epilepsy from the general population Age at follow up, years, mean (SD): Children: 11.7 (8 years) Adults: 30.2 (11 years)	• MRI 1.5-t	 Proportion identified with a clinically relevant abnormality[¥] Proportion identified with a non-epilepsy related abnormality
Byars 2007 Prospective cohort US	N=249 children with a first recognised seizure Age at follow-up, years, mean (SD): 9.6 (2.5)	• MRI 0.5 or 1.5-t	 Proportion identified with a clinically relevant abnormality Proportion identified with a non-epilepsy related abnormality
Coryell 2018 Prospective cohort US	N=714 infants with early life epilepsy Age at seizure onset, months, mean (SD): 11.1 (SD not reported) Age at follow-up, months, mean (SD): 12.7 (SD not reported)	• MRI 1.5 or 3.0-t	 Proportion identified with a clinically relevant abnormality Proportion identified with a non-epilepsy related abnormality
Craven 2012 Retrospective cohort UK	N=2000 young people with focal epilepsy Age at follow-up, years, median (range): 23 (25 to 48)	• MRI 3.0-t	 Proportion identified with a clinically relevant abnormality Proportion identified with a non-epilepsy related abnormality
Das 2013 Cross-sectional India	N=144 infants with epilepsy from the general population Age at seizure onset, years, mean (SD): 2.91 (3.30) Age at follow up, years, mean (SD): 5.87 (4.19)	• MRI 1.5 or 3.0-t	 Proportion identified with a clinically relevant abnormality Proportion identified with a non-epilepsy related abnormality

Study	Population	Intervention	Outcomes
Dirik 2018 Retrospective cohort Cyprus	N=222 infants with newly diagnosed epilepsy Age at seizure onset, months, mean (SD): 48 (SD not reported)	• MRI 1.5 or 3.0-t	 Proportion identified with a clinically relevant abnormality Proportion identified with a non-epilepsy related abnormality
Dura-Trave 2012 Retrospective cohort Spain	N=457 people with epilepsy from the general population Age range at time of diagnosis: 1 month to 15 years	MRI (strength of magnet was not reported)	 Proportion identified with a clinically relevant abnormality Proportion identified with a non-epilepsy related abnormality
Ekici 2013 Retrospective cohort Turkey	N=264 people with epilepsy from the general population Age at follow-up, years, mean (range): 31.3 (18 to 82)	• MRI 3.0-t	 Proportion identified with a clinically relevant abnormality Proportion identified with a non-epilepsy related abnormality
Ferreira 2004 Retrospective cohort Brazil	N=67 adults with focal epilepsy Age at follow-up, years, mean (range): 35 (8 to 76)	• MRI 2.0-t	Proportion identified with a clinically relevant abnormality
Gaillard 2007 Retrospective cohort US	N=38 children with focal epilepsy Age at seizure onset, years, mean (range): 5.8 (0.9 to 11.9)	• MRI 1.5-t	Proportion identified with a clinically relevant abnormality
Griffiths 2005 Retrospective cohort UK	N= 120 young people with focal epilepsy Age at seizure onset, years, median (range): 13 (25 to 38)	• MRI 3.0-t	Proportion identified with a clinically relevant abnormality

Study	Population	Intervention	Outcomes
Hakami 2013 Prospective cohort Australia	N=764 adults with new-onset epilepsy Age at follow-up, years, mean (SD/range): 42.2 (18.8/14.3 to 94.3)	• MRI 1.5 or 3.0-t	 Proportion identified with a clinically relevant abnormality Proportion identified with a non-epilepsy related abnormality
Harini 2018 Retrospective cohort US	N=71 children with infantile spasms Age at seizure onset, years, median (IQR): 6 (IQR not reported)	• MRI 1.5 or 3.0-t	Proportion identified with a clinically relevant abnormality
Hesdorffer 2008 Prospective cohort US	N=159 infants with febrile seizures Age at seizure onset, months (%): <18 months, n=75 (47.2); ≥18 months, n=84 (52.8)	• MRI 1.5-t	 Proportion identified with a clinically relevant abnormality Proportion identified with a non-epilepsy related abnormality
Hnojcikova 2010 Retrospective cohort US	N=28 children with epilepsy from the general population Age at seizure onset, months, mean years (SD): 9.6 (12.7) Age at follow-up, months, mean (SD): 28.8 (17.7)	MRI (strength of magnet was not reported)	Proportion identified with a clinically relevant abnormality
Hsieh 2010 Prospective cohort US	N=182 infants with new onset afebrile seizures At follow-up, all infants were <24 months	• MRI 1.5-t	 Proportion identified with a clinically relevant abnormality Proportion identified with a non-epilepsy related abnormality
Jasim 2018 Cross-sectional Iraq	N=51 people with epilepsy from the general population Age at follow up, mean years (SD): 21.31 (12.75)	• MRI 1.5-t	 Proportion identified with a clinically relevant abnormality

Study	Population	Intervention	Outcomes
Jeniffer 2015 Prospective cohort India	N=64 people with focal seizures At follow-up, all were <18 years old	• MRI 1.5-t	Proportion identified with a clinically relevant abnormality
Koirala 2011 Cross-sectional Nepal	N=160 people with epilepsy from the general population Age at follow-up, years: range was 1 to 82 years old; n=36 (22.5) were ≥16 years old; n=124 (77.5) were >16 years old	• MRI 0.2-t	Proportion identified with a clinically relevant abnormality
Labate 2006 Retrospective cohort Italy	N=101 young people with focal epilepsy Age at seizure onset, years, mean (SD): 22.3 (17.4 years) Age at follow-up, years, mean (SD): 37.3 (17.5)	• MRI 1.5-t	 Proportion identified with a clinically relevant abnormality
Lefkopoulos 2005 Retrospective cohort Greece	N=120 young people with intractable partial seizures Age at follow-up, years, mean (SD): 21 (SD not reported)	• MRI 1.5-t	 Proportion identified with a clinically relevant abnormality
Ma 2019 Retrospective cohort China	N=115 adults with focal epilepsy Age at follow-up, years, mean (SD): 30.8 (12.6)	MRI (strength of magnet not reported)	Proportion identified with a clinically relevant abnormality
Nair 2009 Prospective cohort India	N=41 adults with status epilepticus Age at follow-up, years, mean (range): 35 (1 to 78)	• MRI 1.5-t	Proportion identified with a clinically relevant abnormality

Study	Population	Intervention	Outcomes
Petrou 2007 Retrospective cohort Sweden	N=437 infants with epilepsy from the general population Age at seizure onset, mean months (SD): 14.1 (not reported)	MRI (strength of magnet not reported)	Proportion identified with a clinically relevant abnormality
Rasool 2012 Prospective cohort India	N=157 people with first onset afebrile and complex febrile seizures Age at follow-up, range: 6 months to 14 years old	• MRI 1.5-t	Proportion identified with a clinically relevant abnormality
Santos 2005 Retrospective cohort Brazil	N=100 children with focal epilepsy Age at seizure onset, years, mean (SD): 8.5 (3.1) Age at follow-up, years, mean (SD): 23.9 (9)	MRI (strength of magnet not reported)	Proportion identified with a clinically relevant abnormality
Sinha 2012 Prospective cohort India	N=43 older people with epilepsy Age at seizure onset, years, mean (SD): 68 (7.5)	• MRI 1.5-t	Proportion identified with a clinically relevant abnormality
Solosrungruang 2007 Retrospective cohort Thailand	N=91 adult people with epilepsy from the general population Age at follow-up, years, mean (range): 36.9 (15-85)	• MRI 1.5-t	Proportion identified with a clinically relevant abnormality
Toledo 2013 Prospective cohort Spain	N=161 adults with focal epilepsy Age at follow-up, years, mean (SD): 41.6 (16.3)	• MRI 3.0-t	Proportion identified with a clinically relevant abnormality

Study	Population	Intervention	Outcomes
Wieshmann 2003 Cross-sectional UK	N=332 adults with epilepsy from the general population Age at follow-up, years, mean (SD): 39.7 (14.2)	• MRI 1.5-t	 Proportion identified with a clinically relevant abnormality
Wongladarom 2004 Retrospective cohort Thailand	N=100 children with epilepsy from the general population Age at follow-up, years, mean (SD): 7 (5 months)	• MRI 1.5-t	 Proportion identified with a clinically relevant abnormality Proportion identified with a non-epilepsy related abnormality

IQR: interquartile range; SD: standard deviation

 Δ This study included people with arteriovenous malformations (AVM) only, therefore the proportion identified with vascular abnormalities was 100%. This study was excluded from the vascular abnormalities estimates, but the results have been noted in the evidence table \pm All infections identified in this study were neurocysticercosis, which is a condition endemic to Bhutan, where the study was conducted

See the full evidence tables in appendix D and the forest plots in appendix E.

Summary of the evidence

Epilepsy related abnormalities (clinically relevant abnormalities) detected by MRI

- Very low quality evidence from 24 observational studies assessing N= 6693
 people with epilepsy showed that the overall proportion of people identified by MRI
 with tumour abnormalities was 3% (95% CI, 2 to 4%). The proportion of tumour
 abnormalities identified by MRI in subgroup analyses were as follows:
 - o By age group:
 - Infants (<3 years old at seizure onset): n= 985, 1% (95% CI, 1 to 2%)
 - Children (between 3 and 11 years old at seizure onset): n= 516, 1% (95% CI, 0 to 2%)
 - Young people (between 11 and 25 years old at seizure onset): n= 120, 3% (95% CI, 1 to 8%)
 - Older people (above 65 years old at seizure onset): n= 43, 12% (95% CI, 4 to 25%)
 - By seizure type:
 - People with focal (partial) epilepsy: n= 2660, 4% (95% CI, 2 to 9%)
 - People with genetic (idiopathic) generalised epilepsy: n= 144, 5% (95% CI, 2 to 14%)
 - By MRI strength of magnet:
 - MRI 1.5-t: n= 1080, 4% (95% CI, 2 to 7%)
 - MRI 3-t: n= 3309, 3% (95% CI, 1 to 6%)
 - o By response to treatment:
 - People with a new diagnosis: n= 1556, 1% (95% CI, 0 to 3%)
 - People with existing diagnosis and treatment resistant: n= 454, 5% (95% CI, 2 to 12%)

- People with existing diagnosis and controlled: n= 170, 0% (95% CI, 0 to 2%)
- o By presence/absence of learning disabilities:
 - People without learning disabilities: n= 64, 2% (95% CI, 0 to 8%)
- By previous CT scan:
 - People with a previous CT scan: n = 269, 4% (95% CI, 1 to 13%)
- Very low quality evidence from 25 observational studies assessing N= 7544
 people with epilepsy showed that the overall proportion of people identified by MRI
 with vascular abnormalities was 6% (95% CI, 4 to 8%). The proportion of vascular
 abnormalities identified by MRI in subgroup analyses were as follows:
 - o By age group:
 - Children (between 3 and 11 years old at seizure onset): n= 559, 4% (95% CI, 1 to 18%)
 - Young people (between 11 and 25 years old at seizure onset): n= 240, 7% (95% CI, 4 to 48%)
 - Older people (above 65 years old at seizure onset): n= 43, 30% (95% CI, 17 to 46%)
 - o By seizure type:
 - People with focal (partial) epilepsy: n= 2596, 4% (95% CI, 2 to 8%)
 - People with genetic (idiopathic) generalised epilepsy: n= 60, 8% (95% CI, 4 to 19%)
 - People with West syndrome: n= 73, 21% (95% CI, 13 to 31%)
 - People with Lennox-Gastaut syndrome: n= 1, 0% (95% CI, 0 to 2%)
 - o By MRI strength of magnet:
 - MRI 1.5-t: n=794, 11% (95% CI, 7 to 17%)
 - MRI 3-t: n= 559, 4% (95% CI, 2 to 7%)
 - o By response to treatment:
 - People with a new diagnosis: n=2370, 4% (95% CI, 2 to 9%)
 - People with existing diagnosis and treatment resistant: n= 426, 6% (95% CI, 4 to 9%)
 - People with existing diagnosis and controlled: n= 170, 2% (95% CI, 0 to 5%)
- Very low quality evidence from 37 observational studies assessing N= 8681
 people with epilepsy showed that the overall proportion of people identified by MRI
 with scarring abnormalities was 10% (95% CI, 6 to 16%). The proportion of
 scarring abnormalities identified by MRI in subgroup analyses were as follows:
 - o By age group:
 - Infants (<3 years old at seizure onset): n= 1858, 4% (95% CI, 2 to 9%)
 - Children (between 3 and 11 years old at seizure onset): n= 625, 17% (95% CI, 4 to 49%)
 - Young people (between 11 and 25 years old at seizure onset): n= 341, 21% (95% CI, 10 to 40%)
 - Adults (between 25 and 65 years old at seizure onset): n= 134, 8% (95% CI, 4 to 14%)
 - Older people (above 65 years old at seizure onset): n= 43, 2% (95% CI, 0 to 12%)
 - By seizure type:
 - People with focal (partial) epilepsy: n= 3023, 17% (95% CI, 8 to 31%)

- People with genetic (idiopathic) generalised epilepsy: n= 467, 8% (95% CI, 2 to 32%)
- Those with West syndrome: n= 171, 7% (95% CI, 3 to 15%)
- Those with Lennox-Gastaut syndrome: n=100, 42% (95% CI, 32 to 52%)
- o By MRI strength:
 - MRI 1.5-t: n = 1687, 12% (95% CI, 6 to 23%)
 - MRI 3-t: n= 3045, 15% (95% CI, 10 to 21%)
- O By response to treatment:
 - People with a new diagnosis: n=2576, 7% (95% Cl, 2 to 18%)
 - People with existing diagnosis and treatment resistant: n=574, 20% (95% CI, 6 to 49%)
 - People with existing diagnosis and controlled: n=202, 11% (95% CI, 3 to 35%)
- o By presence/absence of learning disabilities:
 - People without learning disabilities: n= 96, 10% (95% CI, 3 to 26%)
- By previous CT scan:
 - People with a previous CT scan: n= 426, 4% (95% CI, 1 to 13%)
- Very low quality evidence from 31 observational studies assessing N= 8450
 people with epilepsy showed that the overall proportion of people identified by MRI
 with congenital/developmental abnormalities was 10% (95% CI, 7 to 15%). The
 proportion of congenital/developmental abnormalities identified by MRI in
 subgroup analyses was as follows:
 - o By age group:
 - Infants (<3 years old at seizure onset): n=1858, 13% (95% CI, 9 to 19%)
 - Children (between 3 and 11 years old at seizure onset): n= 587, 27% (95% CI, 12 to 48%)
 - Young people (between 11 and 25 years old at seizure onset): n= 240, 9% (95% CI, 2 to 27%)
 - Adults (between 25 and 65 years old at seizure onset): n= 134, 2% (95% CI, 0 to 6%)
 - By seizure type:
 - People with focal (partial) epilepsy: n=2810, 9% (95% CI, 5 to 18%)
 - People with genetic (idiopathic) generalised epilepsy: n=307, 3% (95% CI, 2 to 6%)
 - By syndrome type:
 - Those with West syndrome: n= 73, 41% (95% CI, 30 to 53%)
 - Those with Lennox-Gastaut syndrome: n=137, 15% (95% CI, 10 to 22%)
 - By MRI strength of magnet:
 - MRI 1.5-t: n= 1422, 16% (95% CI, 9 to 26%)
 - MRI 3-t: n=3309, 4% (95% CI, 3 to 7%)
 - By response to treatment:
 - People with a new diagnosis: n=2676, 9% (95% CI, 5 to 15%)
 - People with existing diagnosis and treatment resistant: n=574, 16% (95% CI, 7 to 33%)
 - People with existing diagnosis and controlled: n= 170, 0% (95% CI, 0 to 2%)
 - By presence/absence of learning disabilities:
 - People with learning disabilities: n= 135, 15% (95% CI, 9 to 22%)

- People without learning disabilities: n= 64, 45% (95% CI, 33 to 58%)
- o By previous CT scan:
 - People with a previous CT scan: n= 339, 14% (95% CI, 4 to 37%)
- Very low quality evidence from 19 observational studies assessing N= 5341
 people with epilepsy showed that the overall proportion of people identified by MRI
 with inflammatory/infective/inmumne abnormalities was 4% (95% CI, 2 to 9%).
 The proportion of inflammatory/infective/immune abnormalities identified by MRI in
 subgroup analyses was as follows:
 - o By age group:
 - Infants (<3 years old at seizure onset): n=1477, 1% (95% CI, 1 to 2%)
 - Children (between 3 and 11 years old at seizure onset): n= 559, 2% (95% CI, 1 to 5%)
 - Young people (between 11 and 25 years old at seizure onset): n= 240, 3% (95% CI, 1 to 6%)
 - Older people (above 65 years old at seizure onset): n= 43, 12% (95% CI, 4 to 25%)
 - By seizure type:
 - People with focal (partial) epilepsy: n=2361, 2% (95% CI, 1 to 8%)
 - People with genetic (idiopathic) generalised epilepsy: n=16, 12% (95% CI, 2 to 38%)
 - o By syndrome type:
 - Those with West syndrome: n= 73, 4% (95% CI, 1 to 12%)
 - Those with Lennox-Gastaut syndrome: n= 2, 0% (95% CI, 0 to 2%)
 - o By MRI strength of magnet:
 - MRI 1.5-t: n= 794, 10% (95% CI, 2 to 31%)
 - MRI 3-t: n= 2120, 1% (95% CI, 0 to 3%)
 - O By response to treatment:
 - People with a new diagnosis: n= 1284, 1% (95% CI, 1 to 2%)
 - People with existing diagnosis and treatment resistant: n= 452, 7% (95% CI, 4 to 13%)
 - By previous CT scan:
 - People with a previous CT scan: n= 266, 13% (95% CI, 1 to 82%)
- Very low quality evidence from 9 observational studies assessing N= 4426 people
 with epilepsy showed that the overall proportion of people identified by MRI with
 metabolic/genetic abnormalities was 1% (95% CI, 1 to 3%). The proportion of
 metabolic/genetic abnormalities identified by MRI in subgroup analyses was as
 follows:
 - o By age group:
 - Infants (<3 years old at seizure onset): n= 1477, 1% (95% CI, 0 to 1%)
 - Children (between 3 and 11 years old at seizure onset): n= 388, 4% (95% CI, 2 to 6%)
 - By seizure type:
 - People with focal (partial) epilepsy: n= 2000, 0% (95% CI, 0 to 1%)
 - By syndrome type:
 - Those with Lennox-Gastaut syndrome: n= 135, 7% (95% CI, 3 to 12%)

- o By MRI strenght of magnet:
 - MRI 1.5-t: n=399, 1% (95% CI, 0 to 3%)
 - MRI 3-t: n= 2000, 0% (95% CI, 0 to 1%)
- O By response to treatment:
 - People with a new diagnosis: n= 1284, 2% (95% CI, 1 to 4%)
 - People with existing diagnosis and treatment resistant: n= 217, 0% (95% CI, 0 to 3%)
- By presence/absence of learning disabilities:
 - People without learning disabilities: n= 135, 7% (95% CI, 3 to 12%)
- By previous CT scan:
 - People with a previous CT scan: n= 182, 2% (95% CI, 0 to 5%)

Non-epilepsy related abnormalities detected by MRI

- Very low quality evidence from 20 observational studies assessing N= 6628
 people with epilepsy showed that the overall proportion of people identified by MRI
 with non-epilepsy related abnormalities was 6% (95% CI, 4 to 9%). The
 proportion of non-epilepsy related abnormalities identified by MRI in subgroup
 analyses was as follows:
 - o By age group:
 - Infants (<3 years old at seizure onset): n= 1421, 8% (95% CI, 3 to 18%)
 - Children (between 3 and 11 years old at seizure onset): n= 388, 4% (95% CI, 2 to 6%)
 - Adults (between 25 and 65 years old at seizure onset): n= 134, 1% (95% CI, 0 to 5%)
 - By seizure type:
 - People with focal (partial) epilepsy: n= 2183, 7% (95% CI, 2 to 22%)
 - People with genetic (idiopathic) generalised epilepsy: n= 383, 4% (95% CI, 2 to 10%)
 - By syndrome type:
 - Those with West syndrome: n = 2, 0% (95% CI, 0 to 84%)
 - Those with Lennox-Gastaut syndrome: n= 137, 1% (95% CI, 0 to 5%)
 - By MRI strength of magnet:
 - MRI 1.5-t: n= 688, 10% (95% CI, 5 to 16%)
 - MRI 3-t: n= 2000, 16% (95% CI, 15 to 18%)
 - O By response to treatment:
 - People with a new diagnosis: n= 2733, 6% (95% CI, 3 to 12%)
 - People with existing diagnosis and treatment resistant: n= 311, 1% (95% CI, 0 to 62%)
 - People with existing diagnosis and controlled: n= 202, 5% (95% CI, 1 to 15%)
 - By presence/absence of learning disabilities:
 - People with learning disabilities: n = 135, 1% (95% CI, 0 to 4%)
 - People without learning disabilities: n= 32, 12% (95% CI, 4 to 29%)
 - By previous CT scan:
 - People with a previous CT scan: n= 383, 7% (95% CI, 2 to 19%)

Quality assessment of clinical outcomes included in the evidence review

See the clinical evidence profiles in appendix F.

Economic evidence

Included studies

A single economic search was undertaken for all topics included in the scope of this guideline but no economic studies were identified which were applicable to this review question. See the literature search strategy in appendix B and economic study selection flow chart in appendix G.

Excluded studies

A single economic search was undertaken for all topics included in the scope of this guideline. See supplementary material 2 for details.

Summary of studies included in the economic evidence review

No studies were identified which were applicable to this review question

Economic model

No economic modelling was undertaken for this review because the committee agreed that other topics were higher priorities for economic evaluation.

Summary of the economic evidence

No evidence was identified which was applicable to this review question.

The committee's discussion of the evidence

Interpreting the evidence

The outcomes that matter most

The committee identified two outcomes as relevant for this review question. As part of the critical outcomes, the committee prioritised the proportion identified with a clinically relevant abnormality. Identification of structural brain abnormalities related with epilepsy may inform additional testing, and the need for surgery in people with epilepsy. As part of the important outcomes, the committee prioritised the proportion with a non-epilepsy abnormality. 'Incidental findings' on scans can be a huge source of worry for people. Some of them will have operations or treatment based on these 'incidental' findings because these can be harmful, even when not associated with epilepsy.

The quality of the evidence

The quality of the evidence was assessed with a modified GRADE approach, using the same principles of GRADE for assessing the quality of the evidence, but a different form of presentation as GRADE is not yet available for single-arm prevalence studies.

The quality of the evidence was considered to be very low for most of the outcomes. The domain 'risk of bias' was assessed with the CEBMA checklist, and most studies

were considered to be at very high risk of bias, mainly due to the sampling approaches used and concerns regarding how representative the samples were.

Many of the outcomes were also downgraded due to high levels of imprecision in the estimated proportions.

Other concerns included very high between-study heterogeneity amongst the included studies, for which random effects model was considered. Possible causes for this substantial heterogeneity are believed to be the variability among the included studies characteristics, such as the variety of designs, point along the pathway when MRI was undertaken, or excessive clinical diversity of the individuals included. It was not considered that sensitivity analyses would identify the cause for heterogeneity as excluding a few studies from the analyses on the basis of specific characteristics could add undue emphasis on post-hoc data dependent analysis. Additionally, it was not believed that this will lead to solid results, particularly when it was already established, by committee's informal consensus that the underlying cause of heterogeneity was not due to a single factor.

As a result of the variability between the included studies, some studies appear to be outliers in the meta-analyses conducted; for example Ma 2019, which contributed to the meta-analysis of proportion of tumours abnormalities identified in focal (partial) epilepsy. The lower 95% CI for Ma 2019 is above the upper 95% CI for the pooled estimate. The results reported by Ma 2019 were pre-operative MRI assessments, so it is anticipated that the sample of people included in this study was highly selective.

Outcomes were downgraded for inconsistency, as appropriate, and the committee interpreted the evidence taking these limitations into consideration.

Overall, the committee agreed that the evidence was of insufficient quality as the basis to make recommendations alone and supplemented the information provided by the review with their clinical experience and awareness of the wider literature.

Benefits and harms

Neuroimaging is one of the most common imaging tests in people with 1 or more confirmed seizures. MRI helps identify the cause of epilepsy and provides the information necessary to plan appropriate treatment.

The evidence showed that the yield of clinically relevant abnormalities varied by age. Infants (<3 years old) and children (3 to 11 years old) had higher yield of congenital/developmental abnormalities; children and young people (>11 to 25 years old) had higher yield of scarring abnormalities; and older people (>65 years old) higher yield of inflammatory/infective/immune and vascular abnormalities. These findings are consistent with the clinical experience and expertise of the committee, who emphasised that MRI scanning is particularly important in those who develop epilepsy before the age of 2 or in adulthood. Onset of seizures in these age groups is more frequently associated with an abnormality demonstrable on neuroimaging. However, the committee agreed that an abnormality could be present at any age and agreed to make a recommendation to this effect.

The committee discussed that there are specific conditions in which neuroimaging is not needed routinely because they are not associated with abnormal findings, namely idiopathic generalised epilepsy (IGE) that responds to treatment, or childhood epilepsy with centrotemporal spikes.

Based on their experience and expertise, the committee established that MRI scans should be offered within 6 weeks of referral to avoid undue delays. The committee

could not recommend a specific imaging protocol as this was not formally assessed in the review. However, to avoid ambiguity, the committee decided to recommend that regionally agreed protocols should be followed. From clinical experience and expertise, the committee noted that these should be detailed enough to pick up relevant and subtle abnormalities that may cause epilepsy. The protocol should include 3D imaging datasets, such as suggested in the International League Against Epilepsy (ILAE) recommendations on structural magnetic resonance imaging (https://pubmed.ncbi.nlm.nih.gov/31135062). Where possible, the scan should be performed on a higher magnetic field strength scanner (3T preferred over 1.5T).

There may be some situations where general anaesthetic or sedation may be required in order for the person to undergo neuroimaging. For example, this would be needed in those who find it difficult to lie still for the scan (particularly children aged 3 months to 5 years) or those who are anxious during imaging, so the benefits and risks of the anaesthetic procedure or sedation should be discussed with them. Other alternatives to help people go through the procedure includes various approaches to facilitate the process, such as desensitisation or administration of anxiolytic drugs prior to the procedure. Play therapy may also help children to prepare for and undertake the scan. The committee emphasised that these measures should be tailored to each situation and person.

The use of CT or MRI is associated with possible harm. For example, if a contrast agent is used, there is a risk of allergic reaction to it. For CT, there is the specific risk of radiation exposure, which is related to the dose of radiation and the age of the person (worse at younger age). There is no radiation risk associated with MRI, but this modality may not be suitable for some people the procedure takes longer than a CT scan, and may provoke feelings of claustrophobia in susceptible individuals. Additionally, unlike CT, MRI is also contraindicated in those with some metallic implants, such non-MR conditional pacemakers. The benefits for each procedure have to be balanced against the associated risks.

The committee discussed that in cases where MRI cannot be tolerated, CT should be considered. The main disadvantage of CT as compared to MRI is that CT is less sensitive in detecting subtle abnormalities, especially developmental abnormalities, although it may help identify the cause of an acute symptomatic seizure.

The committee acknowledged that paediatric neuroradiologists within tertiary centres have expertise in reporting children's scans, and their expertise can be sought when there are doubts regarding the relevance of imaging findings on children's or young people's scans or in cases of children or young people with drug resistant epilepsy. The committee explained that interpretation of imaging in children and young people can be challenging due to the complex structural brain changes that take place during child development.

The use of agreed epilepsy protocols should reduce the requirement for repeat scans, saving resources over time. However, the committee noted that there are some situations when a repeat MRI scan may be needed. This includes if the first scan was suboptimal, or was done many years ago (as there has been improvement in neuroimaging with modern scanners and scanning techniques), if new symptoms have appeared, or if surgery is being considered.

Cost effectiveness and resource use

The committee noted that no relevant published economic evaluations had been identified and no additional economic analysis had been undertaken in this area.

In current practice, most people with epilepsy will receive neuroimaging to help identify their cause of epilepsy. Therefore, the committee agreed to make a strong recommendation about offering neuroimaging to people with 1 or more confirmed epileptic seizures, in order to look for an underlying cause and assist in planning appropriate treatment. This reflects current practice, so there will not be substantial impact on use of NHS resources associated with these recommendations. There may be some cost savings from refining the diagnostic pathway and reducing the requirement for repeat investigations.

The committee agreed that there would be minimal impact on resource use in the way the MRI scans are conducted, reported and reviewed, as these recommendations largely reflect current practice.

Finally, the committee discussed the length of time people with epilepsy should be expected to wait for neuroimaging. According to the NHS constitution diagnostic imaging should be undertaken within 6 weeks from the referral. The committee considered this was appropriate.

Recommendations supported by this evidence review

This evidence review supports recommendation section 1.3.1-1.3.7.

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Appendices

Appendix A – Review protocols

Review protocol for review question: What is the yield of relevant abnormalities detected by MRI in people with epilepsy?

Table 3: Review protocol for yield of relevant abnormalities detected by MRI in people with epilepsy

Field	Content
PROSPERO registration number	CRD42019159416
Review title	Magnetic resonance imaging scan to detect relevant abnormalities in people with epilepsy
Review question	What is the yield of relevant abnormalities detected by MRI in people with epilepsy?
	Note: The question has changed from that in the scope, as the committee agreed the accuracy of MRI is known; however determining when MRI should be used is not clear
Objective	The objective of this review is to assess how well magnetic resonance imaging (MRI) performs in detecting brain lesions or other relevant abnormalities in people with epilepsy. Knowing the frequency of these abnormalities, helps clinicians to recognise those people who are most at risk of
	adverse outcomes, and helps to optimise therapeutic options.
Searches	The following databases will be searched: CDSR CENTRAL DARE HTA MEDLINE & MEDLINE In-Process and Other Non-Indexed Citations Embase

Field	Content
	• EMCare
	Searches will be restricted by:
	Date: year 2000 onwards (because of the MRI Technology advances since that year)
	English language studies
	Human studies
Condition or domain being studied	Epilepsy
Population	Inclusion:
	People with 1 or more confirmed epileptic seizures
	Exclusion:
	Newborn babies (under 28 days) with acute symptomatic seizures
Interventions	Magnetic resonance imaging (MRI)
Comparator	Not relevant
Types of study to be	Systematic reviews of observational studies
included	Prospective/ retrospective cohort studies
	Cross-sectional studies
	Note: For further details, and the algorithm in appendix H. Developing NICE guidelines, the manual
Other evaluation oritoria	Note: For further details, see the algorithm in appendix H, Developing NICE guidelines: the manual.
Other exclusion criteria	 Studies with a mixed population (this is, including children, young people and adults with epilepsy and others with a condition different to epilepsy) will be excluded, unless subgroup analysis for epilepsy has been reported.
Context	Recommendations will apply to those receiving care in any healthcare setting (for example, community, primary,
	secondary care) Priority in decision making will be given to identified studies which report data on both MRI and CT as determining
	who should be tested for MRI and/or CT is required when determining the aetiology of epilepsy.
Primary outcomes (critical outcomes)	Proportion identified with a clinically relevant abnormality

Field	Content
Secondary outcomes (important outcomes)	Proportion identified with a non-epilepsy related abnormality
Data extraction (selection and coding)	All references identified by the searches and from other sources will be uploaded into STAR and de-duplicated. Titles and abstracts of the retrieved citations will be screened. The full text of potentially eligible studies will be retrieved and will be assessed in line with the inclusion criteria outlined in the review protocol. Studies that fail to meet the inclusion criteria once the full version has been checked will be excluded at this stage. Each study excluded after checking the full version will be listed, along with the reason for its exclusion. A standardised form will be used to extract data from studies (see Developing NICE guideline: the manual section 6.4) and will include: study setting; study design; study aim; study dates; funding; sample size; participant demographics and baseline characteristics; inclusion and exclusion criteria; details of intervention and control groups; study methodology; recruitment and study completion rates; outcomes and times of measurement; and information for assessment of risk of bias. All data extraction will be quality assured by a senior reviewer. Draft included and excluded studies tables will be circulated to the Topic Group for their comments. Resolution of disputes will be by discussion between the senior reviewer, Topic advisor and Chair. Duplicate screening will not be undertaken for this question.
Risk of bias (quality) assessment	Quality assessment of individual studies will be performed using the following checklists: ROBIS tool for systematic reviews The CEBMA checklist for prevalence data The quality assessment will be performed by one reviewer and this will be quality assessed by a senior reviewer
Strategy for data synthesis	Depending on the availability of the evidence, the findings will be summarised narratively or quantitatively. Data synthesis Data will be extracted from the studies, and where possible, meta-analyses will be conducted using R, version 3.1.2. A fixed effect meta-analysis will be conducted and data will be presented as absolute rates of yield. Heterogeneity

Field	Content
	Heterogeneity in the effect estimates of the individual studies will be assessed using the I ² statistic. I ² values of greater than 50% and 75% will be considered as significant and very significant heterogeneity, respectively.
	In the presence of heterogeneity, sub-group analysis will be conducted:
	according to the risk of bias of individual studies
	study location
	Exact sub-group analysis may vary depending on differences identified within included studies. If heterogeneity cannot be explained using these methods, random effects model will be used. If heterogeneity remains above 75% and cannot be explained by sub-group analysis; reviewers will consider if meta-analysis is appropriate given characteristics of included studies.
	Validity The confidence in the findings across all available evidence will be evaluated for each outcome using an adaptation of the 'Grading of Recommendations Assessment, Development and Evaluation (GRADE) toolbox' developed by the international GRADE working group: http://www.gradeworkinggroup.org/
Analysis of sub-groups	Stratification Results will be presented separately by:
	Age group:Infants (< 3 years old)
	o Children (3 to 11 years old)
	∘ Young people (> 11 to 25 years old)
	o Adults (> 25 to 65 years old)
	o Older people (> 65 years old)
	Seizure type:
	o Focal (partial)
	o Genetic (idiopathic) generalised

Field	Content	
	Syndrome type: Rolandic West Dravet Lennox Gastaut MRI strength of mag Response to treatme New diagnosis Existing diagnosis Existing diagnosis Learning disability (p	and treatment resistant and controlled present/absent)
Type and method of review		Intervention Diagnostic
		Prognostic
		Qualitative
		Epidemiologic
		Service Delivery
		Other (please specify)
Language	English	

Field	Content		
Country	England		
Anticipated or actual start date	16 January 2020		
Anticipated completion date	21 April 2021		
Stage of review at time	Review stage	Started	Completed
of this submission	Preliminary searches	x	X
	Piloting of the study selection process	X	х
	Formal screening of search results against eligibility criteria	X	х
	Data extraction	x	X
	Risk of bias (quality) assessment	x	X
	Data analysis	x	X
Named contact	5a. Named contact National Guideline Alliance 5b. Named contact e-mail epilepsies@nice.org.uk 5c. Organisational affiliation of the review National Institute for Health and Care Excellence (NICE) and National Guideline Alliance		
Review team members	NGA technical team		
Funding sources/sponsor	This systematic review is being completed by the National Guideline Alliance, which is funded by NICE and hosted by the Royal College of Obstetricians and Gynaecologists. NICE funds the National Guideline Alliance to develop guidelines for those working in the NHS, public health, and social care in England.		
Conflicts of interest	All guideline committee members and review team and expert witnesses) mufor declaring and dealing with conflicts	ist declare any potential conflicts of int	erest in line with NICE's code of practice

Field	Content
	declared publicly at the start of each guideline committee meeting. Before each meeting, any potential conflicts of interest will be considered by the guideline committee Chair and a senior member of the development team. Any decisions to exclude a person from all or part of a meeting will be documented. Any changes to a member's declaration of interests will be recorded in the minutes of the meeting. Declarations of interests will be published with the final guideline.
Collaborators	Development of this systematic review will be overseen by an advisory committee who will use the review to inform the development of evidence-based recommendations in line with section 3 of Developing NICE guidelines: the manual . Members of the guideline committee are available on the NICE website: https://www.nice.org.uk/guidance/indevelopment/gid-ng10112
URL for published protocol	https://www.crd.york.ac.uk/prospero/display_record.php?ID=CRD42019159416
Dissemination plans	NICE may use a range of different methods to raise awareness of the guideline. These include standard approaches such as: notifying registered stakeholders of publication publicising the guideline through NICE's newsletter and alerts issuing a press release or briefing as appropriate, posting news articles on the NICE website, using social media channels, and publicising the guideline within NICE.
Keywords	Genetic testing, yield, management, epilepsy
Details of existing review of same topic by same authors	Not applicable
Additional information	Not applicable
Details of final publication	www.nice.org.uk

CDSR: Cochrane Database of Systematic Reviews; CEBMA; center for evidence-based management; CENTRAL: Cochrane Central Register of Controlled Trials; GRADE: Grading of Recommendations Assessment, Development and Evaluation; HTA: Health Technology Assessment; MID: minimal important difference; NICE: National Institute for Health and Care Excellence; RCT: Randomised Controlled Trial; RoB: Risk of Bias;; ROBIS: risk of bias in systematic reviews; SD: Standard Deviation

Appendix B – Literature search strategies

Literature search strategies for review question: What is the yield of relevant abnormalities detected by MRI in people with epilepsy?

Clinical

Database(s): EMCare, MEDLINE and Embase (Multifile) - OVID

EMCare 1995 to November 25, 2019; Embase Classic+Embase 1947 to 2019 November 25; Ovid MEDLINE(R) and Epub Ahead of Print, In-Process & Other Non-Indexed Citations and Daily 2019 November 25, 2019

Date of last search: 25 November 2019

Multifile database codes: emcr=EMCare; emczd=Embase Classic+Embase; ppez= MEDLINE(R) and Epub Ahead of Print, In-Process & Other Non-Indexed Citations and Daily

exp epilepsy/ use ppez, emczd, emcr or epilep*.ti,ab. (((absence or astatic or atonic or tonic clonic) adj2 (seizure* or spasm*)) or (benign adj2 (childhood or neonatal or pediatric or paediatric) adj2 (convulsion* or seizure* or spasm*)) or (benign adj3 convulsion* adj2 centrotemporal adj2 spike*) or ((centralopathic or centrotemporal or temporal-central focal) adj (convulsion* or seizure*) or continous spike wave of slow sleep or doose* or dravet or ((early or infantile) adj2 mycolonic adj2 encephalopath*) or ((flexor or infantile or neonatal) adj2 (seizure* or spasm*)) or hypsarrhythmia* or infant* spasm* or ((flexor or infantile or neonatal) adj2 (seizure* or spasm*)) or hypsarrhythmia* or infant* spasm* or ((flexor or spack nife or lightening or nodding or salaam) adj (attack* or convulsion* or seizure* or spasm*)) or (mycolonic adj3 (seizure* or spasm*)) or (mycolonic adj3 (seizure* or spasm*)) or (mycolonic adj3 (seizure* or spasm*)) or (propulsive petit mal or spasm in*1 flexion or spasmus nutans or west syndrome*),ti,ab. (bcects or bects or brec or cects or lgs or mae or smei),ti,ab. (bcects or bects or brec or cects or lgs or mae or smei),ti,ab. (bcects or bects or brec or cects or lgs or mae or smei),ti,ab. 4 and 5 exp magnetic resonance imaging/ use ppez or exp nuclear magnetic resonance imaging/ use emczd, emcr (magnetic resonance or mri or mrs or nmr* or ((magneti* or mr or nuclear or nm) adj2 (angiogra* or elastogra* or examin* or imag* or scan* or spectroscop* or tomogra* or tomoangiogra*))),ti,ab. or/7-8 brain injuries/ use ppez or exp brain injury/ use emczd, emcr or ((brain* or cerebral) adj2 (abnormal* or damage or lesion* or malformation*)),ti,ab. exp hemorrhage/ or (bleeding or (blood adj (effusion or loss)) or ha?morrhag* or he?morrhag*),ti,ab. exp hemorrhage/ or (bleeding or (blood adj (effusion or loss)) or ha?morrhag* or he?morrhag*),ti,ab. exp hydrocephalus/ use ppez, emczd, emcr or (anaeductal stenos?s or cerebral ventriculomegal* or hydrocephalus/ use ppez, emczd, emcr o	#	searches
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swelling).ti,ab. exp brain neoplasms/ use ppez or meningioma/ use ppez, emczd, emcr or exp brain tumor/ use emczd, emcr (((brain or cerebral or intracranial or meninges or midline) adj2 (cancer* or metastases or neoplasm* or tumor* or tumour*)) or cerebroma* or mening?oma*).ti,ab. posterior leukoencephalopathy syndrome/ use ppez or posterior reversible encephalopathy syndrome/ use emczd, emcr or ((posterio?r adj (leukoencephalopath* or leuko encephalopath*)) or (posterio?r adj2 reversible encephalopath*) or pres or rpls).ti,ab.	16	hydrocephal*).ti,ab.
emcr ((((brain or cerebral or intracranial or meninges or midline) adj2 (cancer* or metastases or neoplasm* or tumor* or tumour*)) or cerebroma* or mening?oma*).ti,ab. posterior leukoencephalopathy syndrome/ use ppez or posterior reversible encephalopathy syndrome/ use emczd, emcr or ((posterio?r adj (leukoencephalopath* or leuko encephalopath*)) or (posterio?r adj2 reversible encephalopath*) or pres or rpls).ti,ab.	17	swelling).ti,ab.
tumor* or tumour*)) or cerebroma* or mening?oma*).ti,ab. 20 posterior leukoencephalopathy syndrome/ use ppez or posterior reversible encephalopathy syndrome/ use emczd, emcr or ((posterio?r adj (leukoencephalopath* or leuko encephalopath*)) or (posterio?r adj2 reversible encephalopath*) or pres or rpls).ti,ab.	18	emcr
use emczd, emcr or ((posterio?r adj (leukoencephalopath* or leuko encephalopath*)) or (posterio?r adj2 reversible encephalopath*) or pres or rpls).ti,ab.	19	tumor* or tumour*)) or cerebroma* or mening?oma*).ti,ab.
exp vasculitis/ use ppez, emczd, emcr or (angiitis or vasculiti*).ti,ab.	20	use emczd, emcr or ((posterio?r adj (leukoencephalopath* or leuko encephalopath*)) or (posterio?r adj2 reversible encephalopath*) or pres or rpls).ti,ab.
	21	exp vasculitis/ use ppez, emczd, emcr or (angiitis or vasculiti*).ti,ab.

segrons segron		
cerebral venous sinus thrombosis or cvst), it.ab. exp ciatrix/ use ppez or exp scarl ves emczd, emcr or (cicatri?ation or scar¹ to r scarring), it.ab. gliosis/ use ppez, emczd, emcr or (glios?s or gliomatosis or microgliosis), it.ab. (hippocampus and sclerosis), sh. or (hippocampal or ammon horn or hippocampus or incisural or mesial temporal or pararhinal) adj sclerosis), it.ab. ulegrita it.ab. exp demyelinating diseases/ use ppez or exp demyelinating disease/ use emczd, emcr or (demyelinating or (demyelinating adj2 (disorder¹ or disease*))), it.ab. exp finalformations of cortical development? use ppez or exp cortical dysplasia/ use emczd, emcr or ((florain cortext or cortical) adj2 (disorder¹ or development malformation*) adj2 cortical development), it.ab. exp neurocutaneous syndromes/ use ppez or phakomatosis/ use emczd, emcr or ((neurocutaneous adj (disorder² or syndrome²)) or phakomatosis/ use emczd, emcror ((neurocutaneous adj (disorder² or syndrome²)) or phakomatosis/ use emczd, emcror ((neurocutaneous adj (disorder² or syndrome²)) or phakomatosis/ use emczd, emcror ((neurocutaneous adj (disorder² or syndrome²)) or phakomatosis/ use emczd, emcror ((neurocutaneous adj (disorder² or syndrome²)) or phakomatosis/ use emczd, emcror (inlinbic encephaliti²) it.ab. 13 "infection/ use ppez or infection* it.ab. 24 "infection/ use ppez or infection* it.ab. 25 exp "orogenital disorders or glycosylation?" use ppez or exp "congenital disorders or glycosylation?" use emczd, emcr (carbohydrate deficient glycoprotein syndrome² or cdg syndrome² or (congenital disorders adj2' glycosylation) or glycanosis cdg or (carbohydrate deficient adj (glycoprotein disorders or inbon emcro*)), it.ab. 26 exp micochondrial diseases/ use ppez or exp lysosome storage disease/ use emczd, emcr or ((micochondrial adj (deficien*) or or exp	#	searches
gliosis/ use ppez, emczd, emcr or (glios?s or gliomatosis or microgliosis), il.a.b. (hippocampus and sclerosis) sh. or (hippocampal or ammon horn or hippocampus or incisural or mesial temporal or pararhinal) adj sclerosis), il.a.b. 27 exp demyelinating diseases/ use ppez or exp demyelinating disease/ use emczd, emcr or (demyelination or or (disorder))) il.a.b. 28 exp "malformations of cortical development"). Il.a.b. 29 exp encorcutaneous syndromes/ use ppez or or phakomatosis/ use emczd, emcr or ((heurocutaneous adj (disorder or syndrome)) or phakomatosis/ luse emczd, emcr or ((heurocutaneous adj (disorder or syndrome)) or phakomatosis/ luse becap hallis/ use ppez or paraneoplastic neuropahilis/ use ppez, emczd, emcr or (allerige adj (leukoencephalipsh*) or leuko encephalisi/ eneuropahilis/ use ppez, emczd, emcr or (sellergic adj (leukoencephalipsh*) or leuko encephalopah*)) or encephalisi/ use ppez or exp "congenital disorders of glycosylation?" use emczd, emcr or (congenital disorders adj.2 glycosylation) or glycanosis cdg or (carbohydrate deficient adj (glycoprotein disorders adj.2 glycosylation) or glycanosis cdg or (carbohydrate deficient adj (glycoprotein disorders or inborn error")), il., ab. 30 exp lysosomal storage diseases/ use ppez or exp lysosome storage disease/ use emczd, emcr or (lysosomal adj (enzyme or storage) adj (diseaser or disorder*)), il., ab. 31 exp (ysosomal storage) adj (diseaser or disorder*)), il., ab. 32 exp intochondrial diseases/ use ppez or exp lysosome storage diseaser/ use emczd, emcr or (intochondrial disorders adj. adj.) (adj.) (adj		(cerebral venous sinus thrombosis or cvst).ti,ab.
(hippocampus and sclerosis) sh. or (hippocampus or ammon horn or hippocampus or incisural or mesial temporal or parathnial) alg sclerosis).ti, ab. degyris ti, ab. exp demyelinating diseases/ use ppez or exp demyelinating disease/ use emczd, emcr or (demyelination or (demyelinating adj2 (disorder* or disease*))).ti, ab. exp malformations of cortical development) ti, ab. malformation*) adj2 cortical adj2 (dispalsais*) or development malformation*)) or ((abnormal* or malformation*) adj2 cortical adj2 (dispalsais*) or development) ti, ab. exp neurocutaneous syndromes/ use ppez or phakomatosis/ use emczd, emcr or ((flororationation*)) adj2 cortical adj2 (dispalsais*) or development); ti, ab. exp neurocutaneous syndromes/ use ppez or phakomatosis/ use emczd, emcr or ((flororationation*)) or encephalitis/ use emczd, emcr or ((allergic adj (leukoencephalitis/ use ppez or paraneoplastic neuropathy) use emczd, emcr or ((allergic adj (leukoencephalitis/ use ppez or exp* (encephalitis) use emczd, emcr or ((allergic adj (leukoencephalitis/ use ppez or exp* or encephalitis) use emczd, emcr or (earborydrate deficient) adja2 exp* (caphorydrate deficient) glycosylation*) or glycanosis cag or (carborydrate deficient adj (glycoprotein disorders or inborn error*)), ti, ab. leukodystrophy* sh. or ((leucodystroph* or metabolic leucoencephalicpa* or very long chain) adj3 deficien*), ti, ab. exp lysosomal storage diseases/ use ppez or exp plysosome storage disease/ use emczd, emcr or (lysosomal adj (enzyme or storage) adj (disease* or disorder*)), ti, ab. exp mitochondrial diseases/ use ppez or exp plysosome storage disease/ use emczd, emcr or (lysosomal adj (enzyme or storage) adj (disease* or disorder*)), ti, ab. exp disorder*) disp* (allergic adja2) adja3 (deficien*) or disease* or disorder*), ti, ab. (allergic adja4 exp disorder*) adja4 (deficien*) or disease* or disorder*), ti, ab. (allergic adja4 exp disorder*), ti, ab. (allergic adja4 exp disorder*), ti, ab. (allergic adja4 exp disorder*), ti, ab. (allergic adja4		
mesial temporal or pararhinal) adj sclerosis Jtl,ab. 27 ulegyria, ti,ab. 28 exp demyelinating diseases/ use ppez or exp demyelinating disease/ use emczd, emcr or (demyelination or (demyelination adj (disorder* or disease*))), ti,ab. 28 exp "malformations of cortical development?" use ppez or exp cortical dysplasia/ use emczd, emcr or (((brain cortex or cortical) adj2 (dysplasia*) or development malformation*)) or ((abnormal* or malformation*) adj2 cortical development), ti,ab. 29 exp neurocutaneous syndromes/ use ppez or phakomatosis/ use emczd, emcr or ((neurocutaneous adj (disorder* or syndromes/) use ppez or phakomatosis/ use emczd, emcr or ((neurocutaneous adj (disorder* or syndrome*)) or encephalitis/ use ppez, emczd, emcr or (limbic encephalitis/ use ppez or paraneoplastic neuropathy/ use emczd, emcr or limbic encephalitis/ ti,ab. 30 exp encephalitis/ use ppez, emczd, emcr or (limbic encephalitis/ use ppez or paraneoplastic neuropathy/ use emczd, emcr or limbic encephaliti*), ti,ab. 31 "infection/ use ppez or infection*, tij,ab. 32 exp "congenital disorders of glycosylation/" use ppez or exp "congenital disorder of glycosylation/" use emczd, emcr or (carbohydrate deficient glycoprotein syndrome* or cod syndrome* or (congenital disorders adj2 glycosylation) or glycanosis cdg or (carbohydrate deficient adj (glycoprotein disorders or inborn error*)), ti,ab. 32 exp (syspanital) or glycanosis cdg or (carbohydrate deficient adj (glycoprotein disorders or inborn error*)), ti,ab. 33 exp (syspanital) or glycanosis cdg or (carbohydrate deficient adj (glycoprotein disorders or inborn error*)), ti,ab. 34 elukodystrophy*, sh. or ((leucodystroph* or metabolic leucoencephalopa* or very long chain) adj3 deficien*), ti,ab. 35 exp mitochondrial diseases/ use ppez or exp lysosome storage diseases/ use emczd, emcr or ((ysosomal adj (enzyme or storage) adj (disease* or disorder*)), ti,ab. 36 exp mitochondrial diseases/ use ppez or exp lysosome storage diseases/ use emczd, emcr or ((mitochondria) adj (deficien*) or		
exp demyelination of (demyelination gal (disorder or disease*)), il, ab. exp "malformations of cortical development" use ppez or exp cortical dysplasia' use emczd, emcr or ((brain cortext or cortical) adj2 (disporder or disease*)), il, ab. exp neurocutaneous syndromes' use ppez or phakomatosis' use emczd, emcr or ((neurocutaneous adja (disorder or syndrome*) use ppez or phakomatosis') use emczd, emcr or ((neurocutaneous adja (disorder or syndrome*) use ppez or phakomatosis') use emczd, emcr or ((neurocutaneous adja (disorder or syndrome*) use ppez or phakomatosis') use emczd, emcr or ((neurocutaneous adja (disorder or syndrome*) or phakomatosis'), it ab. exp encephalitis' use ppez, emczd, emcr or (imbic encephalitis' use ppez or praneoplastic neuropathy' use emczd, emcr or (imbic encephaliti*), it ab. "infection' use ppez or infection*; it ab. exp "congenital disorders of glycosylation" use ppez or exp "congenital disorder of glycosylation" use emczd, emcr ((arbohydrate deficient glycoprotein syndrome* or cod syndrome* or (congenital disorders adj2 glycosylation) or glycanosis odg or (carbohydrate deficient adj (glycoprotein disorders or inborn error*)), it, ab. leukodystrophy*. sh. or ((leucodystroph* or metabolic leucoencephalopa* or very long chain) adj3 deficien*), it, ab. exp lysosomal storage diseases' use ppez or exp lysosome storage diseases' use emczd, emcr or ((lysosomal adj (enzyme or storage) adj (disease* or disorder*)), it, ab. exp mitochondrial diseases' use ppez or exp "disorders of mitochondrial functions*/ use emczd, emcr or ((mitochondrial adj (deficien*) or disease* or disorder*)) or mitochondrial path or ((election transport chain or oxidative phosphorylation or respiratory chain a) adj2 (deficien* or (election transport chain or oxidative phosphorylation or respiratory chain a) adj2 (deficien*) or (election transport chain or oxidative phosphorylation or respiratory chain a) adj2 (deficien*) or (election transport chain oxidate and deficiency), it, ab. (disorder* adj3 (amino adid*		mesial temporal or pararhinal) adj sclerosis).ti,ab.
exp "malformations of cortical development" use pepz or exp cortical dysplasia/ use emczd, emcr or ((thrain cortext or cortical) adj2 (dysplasia* or development malformation*)) or ((abnormal* or malformation*) adj2 cortical development)).ti, ab. exp neurocutaneous syndromes* use ppez or phakomatosis/ use emczd, emcr or ((neurocutaneous adj (disorder* or syndrome*)) or phakoma* or phacomatos*).ti, ab. exp neurocutaneous syndromes* use ppez or phakomatosis/ use emczd, emcr or ((neurocutaneous adj (disorder* or syndrome*)) or phakoma* or phacomatos*).ti, ab. exp recephaltis* use pez, emczd, emcr or (latilegic adj (leukoencephalopath* or leuko encephality)) or encephaltis* or limbic encephaltis* use ppez or paraneoplastic neuropathy* use emczd, emcr or ((allergic adj (leukoencephalopath* or leuko encephaltis*)) or encephaltis* or limbic encephaltis* use ppez or infection* (i.i.ab. exp 'congenital disorders of glycosylation*)* use ppez or exp "congenital disorder of glycosylation* use emczd, emcr (achohydrate deficient glycoprotein syndrome* or cdg syndrome* or (congenital disorders adj2 glycosylation) or glycanosis cdg or (carbohydrate deficient adj (glycoprotein disorders or inborn error*)); it. ab. leukodystrophy*.sh. or ((leucodystroph* or metabolic leucoencephalopa* or very long chain) adj3 deficien*); it. ab. exp lysosomal storage diseases/ use ppez or exp 'disorder*)) it. ab. exp lysosomal storage diseases/ use ppez or exp 'disorder*)) in mitochondrial functions*/ use emczd, emcr or ((mitochondrial adj (deficien* or disease*) or respiratory chain or oxidative phosphorylation or respiratory chain) adj2 (deficien* or disease* or disorder*)), ab. amino acid metabolism, inborn errors/ use ppez or "disorders of amino acid and protein metabolism*/ use emczd, emcr or (organic adj (acidemia or adicidar*)), ab. ((disorder* adj3 (arnino acid* or protein*) adj3 metaboli*) or (phenyl ketonuria* or phenylketonuria* or tyrosimemia* or homocystinuria* or non-ketotic hypertypicienia* or maple syrup urine disease) or (amino		
(((brain cortext or cortical) adi/2 (dyspiasia* or development maiformation*) or ((abnormal* or maiformation*) adi/2 cortical development)), tiab. generoculaneous syndromes/ use ppez or phakomatosis/ use emczd, emcr or ((neuroculaneous adi/disorder* or syndrome*)) or phakomatos*), tiab. exp encephalitis/ use ppez, emczd, emcr or limbic encephalitis/ use ppez or paraneoplastic neuropathy/ use emczd, emcr or ((allerigic adi/ (leukoencephalopath* or leuko encephalopath*)) or encephalitir or limbic encephalitir/ use emczd, emcr or (congenital disorders of glycosylation*)/ use emczd, emcr infection/ use ppez or infection*.ti,ab. gylocosylation) or glycanosis cdg or (carbohydrate deficient adi/ (glycoprotein disorders or inborn error*)), ti, ab. leukodystrophy*, sh. or ((leucodystroph* or metabolic leucoencephalopa* or very long chain) adi/ deficien*), ti, ab. exp lysosomal storage diseases/ use ppez or exp lysosome storage disease/ use emczd, emcr or ((ysosomal adi/ (enzymer or storage) adi/ (disease* or disorder*)), ti, ab. exp mitochondrial diseases/ use ppez or exp sisorders of mitochondrial functions*/ use emczd, emcr or ((mitochondrial adi/ (deficien*) or disease* or disorder*)) or mitochondriopath* or ((electron transport chain or oxidative phosphorylation or respiratory chain) adi/2 (deficien* or disease* or disorder*)), ti, ab. molybdenum cofactor deficiency / use emczd, emcr or (molybdenum adi/ (co factor or cofactor)) adi/ deficiency), ti, ab. (glucose transporter*, sh. and deficiency), hw. or ((sulfite adi/2) oxidase adi/2 deficiency) or isod), ti, ab. (glucose transporter*, sh. and deficien*) or inseprity or maiformation*, hw. exp epilepsy/di or diagnos* sh. or (diagnos* or detect* or identif* or indicat* or reveal* or ((epilepsy or secure) ad protocor*) or rolesion* or maiformation*), ti, ab. or maiformation*, hw.) use ppez or		(demyelination or (demyelinating adj2 (disorder* or disease*))).ti,ab.
(disorder* or syndrome*)) or phakoma* or phacomatos*),ti,ab. expenephalitis* use ppez, emczd, emc or limbic encephalitis* use ppez or paraneoplastic neuropathy/ use emczd, emcr or ((allergic adj (leukoencephalopath* or leuko encephalopath*)) or encephalitin* or limbic encephaliti*, ti,ab. infection/ use ppez or infection*; ti,ab. exp "congenital disorders of glycosylation*/ use ppez or exp "congenital disorder of glycosylation*/ use emczd, emcr (carbohydrate deficient glycoprotein syndrome* or cdg syndrome* or (congenital disorders adj2 glycosylation) or glycanosis cdg or (carbohydrate deficient adj (glycoprotein disorders or inborn error*))).ti,ab. leukodystrophy*,sh. or ((leucodystroph* or metabolic leucoencephalopa* or very long chain) adj3 deficien*) ti,ab. exp lysosomal at glerayme or storage) adj (disease* or disorder*)),ti,ab. exp lysosomal at glerayme or storage) adj (disease* or disorder*)),ti,ab. exp mitochondrial adj (deficien* or disease*) or disorder*) or mitochondrial functions*/ use emczd, emcr or ((mitochondrial adj (deficien* or disease*) or disorder*)) or mitochondrial functions*/ use emczd, emcr or (organic adj (addemia or addivar*)),ti,ab. molybdenum cofactor deficiency / use emczd, emcr or (molybdenum cofactor deficiency) i.use emczd, emcr or (organic adj (addemia or addivar*)),ti,ab. (disorder* adj3 (amino acid* or protein*) adj3 metabol*) or (phenyl ketonuria* or phenylketonuria* or tyrosimemia* or homocystinuria* or non-ketotic hyperglycinemia* or maple syrup urine disease) or tyrosimemia* or homocystinuria* or non-ketotic hyperglycinemia* or maple syrup urine disease) or (amino acid metablism adj3 inborn error*)),ti,ab. (glucose transporter*,sh. and deficien*,h.v.) or ((glucose transporter adj3 deficien*) or glut1),ti,ab. (glucose transporter*,sh. and deficien*,h.v.) or ((glucose transporter*) or maple or sectional or (diagnos*)		(((brain cortext or cortical) adj2 (dysplasia* or development malformation*)) or ((abnormal* or malformation*) adj2 cortical development)).ti,ab.
neuropathy! use emczd, emcr or ((allergic adj (leukoencephalopath* or leuko encephality) or encephality.it.pib. infection/ use ppez or infection*.ti,ab. infection/ use ppez or infection*.ti,ab. exp "congenital disorders of glycosylation*/ use ppez or exp "congenital disorder of glycosylation"/ use emczd, emcr (carbohydrate deficient glycoprotein syndrome* or cdg syndrome* or (congenital disorders adj2 glycosylation) or glycanosis cdg or (carbohydrate deficient adj (glycoprotein disorders or inborn error")).it.ab. leukodystrophy*.sh. or ((leucodystroph* or metabolic leucoencephalopa* or very long chain) adj3 deficien*).it.jab. exp lysosomal storage diseases/ use ppez or exp lysosome storage disease/ use emczd, emcr or (lysosomal adj (enzyme or storage) adj (disease* or disorder*)).it.jab. exp mitochondrial diseases/ use ppez or exp "disorders of mitochondrial functions" use emczd, emcr or ((mitochondrial diseases/ use ppez or disorder*) or mitochondrial functions" use emczd, emcr or ((mitochondrial diseases) use ppez or disorders of mitochondrial functions" use emczd, emcr or (congenicad disease) (deficien* or disease* or disorder*)) amino acid and protein metabolism. inborn errors/ use ppez or "disorders of amino acid and protein metabolism" use emczd, emcr or (organica adj (acidemia or aciduria*)).it.jab. molybdenum cofactor deficiency / use emczd, emcr or (molybdenum adj (co factor or cofactor) adj deficiency), it.jab. (glucose transporter*, sh. and deficiency).it.yab. (glucose transporter*, sh. and deficiency).it.yab. (glucose transporter*, sh. and deficient*).it.yab. (glucose transporter*, sh. and deficient*).it.yab. or ((glucose transporter adj3 deficien*) or glut1).ti,ab. (glucose transporter*, sh. and deficient*, or (diagnos* or detect* or identif* or indicat* or reveal* or ((epilepsy or secure) adj protocol*) or yield*).ti,ab. 6 and 9 and (magnation*).ti,ab. or (diagnos* or elastogra* or examin*) adj3 (abnormal* or lesion* or malformation*).ti,ab. or malformation*).ti,ab. or malformation*)	29	(disorder* or syndrome*)) or phakoma* or phacomatos*).ti,ab.
exp "congenital disorders of glycosylation"/ use ppez or exp "congenital disorder of glycosylation"/ use emczd, emcr (carbohydrate deficient glycoprotein syndrome* or cdg syndrome* or (congenital disorders adj2 glycosylation) or glycanosis cdg or (carbohydrate deficient adj (glycoprotein disorders or inborn erro*)).ti, ab. leukodystrophy* sh, or ((leucodystroph* or metabolic leucoencephalopa* or very long chain) adj3 deficien*), ti, ab. exp lysosomal storage diseases/ use ppez or exp lysosome storage disease/ use emczd, emcr or (lysosomal adj (enzyme or storage) adj (disease*) or disorder*)), ti, ab. exp mitochondrial diseases/ use ppez or exp "disorders of mitochondrial functions"/ use emczd, emcr or ((mitochondrial adj (deficien*) or disease* or disorder*)) or mitochondriopath* or ((electron transport chain or oxidative phosphorylation or respiratory chain) adj2 (deficien*) or disorder*)), ti, ab. molybdenum cofactor deficiency / use emczd, emcr or (molybdenum adj (co factor or cofactor) adj deficiency), ti, ab. ((disorder* adj3 (amino acid* or protein*) adj3 metaboli*) or (phenyl ketonuria* or phenylketonuria* or tyrosinemia* or homocystinuria* or non-ketotic hyperglycinemia* or maple syrup urine disease) or (amino acid metabolism adj3 inborn error*), ti, ab. ((glucose transporter*, sh. and deficien*, hw.) or ((glucose transporter adj3 deficien*) or glut1), ti, ab. ((or/10-41) or (abnormal* or lesion* or malformation*), ti, ab. or malformation*, hw. exp epilepsyldi or diagnos*.sh. or (diagnos* or detect* or identif* or indicat* or reveal* or ((epilepsy or seizure) adj protocol*) or tomogra* or elastogra* or examin*) adj3 (abnormal* or lesion* or malformation*). ti, ab. 6 and 9 and 42 and 43 6 and 9 and ((magnetic resonance or mri or mrs or nm* or angiogra* or tomoangiogra* or imag* or scan* or spectroscop* or tomogra* or elastogra* or examin*) adj3 (abnormal* or lesion* or malformation*), ti, ab. or malformation*), ti, ab. or malformation*, ti, ab. or malformation*, ti, ab. or or or or or or or or		neuropathy/ use emczd, emcr or ((allergic adj (leukoencephalopath* or leuko encephalopath*)) or encephaliti* or limbic encephalit*).ti,ab.
emczd, emcr (carbohydrate deficient glycoprotein syndrome* or cdg syndrome* or (congenital disorders adj2 glycosylation) or glycanosis cdg or (carbohydrate deficient adj (glycoprotein disorders or inborn emror*)), it, ab. leukodystrophy* sh, or ((leucodystroph* or metabolic leucoencephalopa* or very long chain) adj3 deficien*), it, ab. sex plysosomal storage diseases/ use ppez or exp lysosome storage disease/ use emczd, emcr or (lysosomal adj (enzyme or storage) adj (disease* or disorder*)), it, ab. exp mitochondrial diseases/ use ppez or exp "disorders of mitochondrial functions"/ use emczd, emcr or (mitochondrial dijedeficien* or disorder*) or mitochondriopath* or ((electron transport chain or oxidative phosphorylation or respiratory chain) adj2 (deficien* or disease* or disorder*))), it, ab. amino acid metabolism, inborn errors' use ppez or disorders of amino acid and protein metabolism?' use emczd, emcr or (organic adj (acidemia or aciduria*)), it, ab. molybdenum cofactor deficiency / use emczd, emcr or (molybdenum adj (co factor or cofactor) adj deficiency), it, ab. ((disorder* adj3 (amino acid* or protein*) adj3 metaboli*) or (phenyl ketonuria* or phenylketonuria* or tyrosimemia* or homocysthuria* or non-ketotic hyberglycinemia* or maple syrup urine disease) or (amino acid metablism adj3 inborn error*)), it, ab. (glucose transporter*.sh. and deficien*.hw.) or ((glucose transporter adj3 deficien*) or glut1), it, ab. (glucose transporter*.sh. and deficien*.hw.) or ((glucose transporter adj3 deficien*) or (lepilepsy or seizure) adj protocol*) or yield*), it, ab. 6 and 9 and ((magnetic resonance or mri or mrs or nmr* or angiogra* or tomoangiogra* or imag* or seartor or spectroscop* or tomogra* or elastogra* or examin*) adj3 (abnormal* or lesion* or malformation*), it, ab. 6 and 9 and (exp case control studies/ or exp cohort studies/) or longitudinal or (observational adj (study or studies)) or retrospective), it, ab.) and ((abnormal* or lesion* or malformation*), it, ab. or malformation*, it, ab. o		
glycosylation) or glycanosis cdg or (carbohydrate deficient adj (glycoprotein disorders or inborn error*)).it.ab. 34 leukodystrophy*s.h. or ((leucodystroph* or metabolic leucoencephalopa* or very long chain) adj3 deficien*).it.ab. 35 exp lyososmal storage diseases/ use ppez or exp lysosome storage disease/ use emczd, emcr or (lysosomal adj (enzyme or storage) adj (disease* or disorder*)).it.ab. 36 exp mitochondrial diseases/ use ppez or exp "disorders of mitochondrial functions"/ use emczd, emcr or ((mitochondrial adj (deficien* or disease* or disorder*)) or mitochondriopath* or ((electron transport chain or oxidative phosphorylation or respiratory chain) adj2 (deficien* or disease* or disorder*)).it.jab. 37 amino acid metabolism, inborn errors/ use ppez or "disorders of amino acid and protein metabolism"/ use emczd, emcr or (organic adj (acidemia or aciduria*)).it.jab. 38 molybdenum cofactor deficiency / use emczd, emcr or (molybdenum adj (co factor or cofactor) adj deficiency), ti.jab. 39 ((sulfite oxidase and deficiency), hw. or ((sulfite adj2 oxidase adj2 deficiency) or isod).ti.jab. 40 ((disorder* adj3 (amino acid* or protein*) adj3 metaboli*) or (phenyl ketonuria* or phenylketonuria* or tyrosinemia* or homocystinuria* or non-ketotic hyperglycinemia* or maple syrup urine disease) or (amino acid metablism adj3 inborn error*)).ti,ab. 41 (glucose transporter* sh. and deficien*, hw.) or ((glucose transporter adj3 deficien*) or glut1).ti,ab. 42 (or/10-41) or (abnormal* or lesion* or malformation*).ti,ab. or malformation* hw. 43 exp epilepsyldi or diagnos* sh. or (diagnos* or detect* or identif* or indicat* or reveal* or ((epilepsy or seizure) adj protocol*) or yield*).ti,ab. 44 6 and 9 and 42 and 43 45 6 and 9 and (magnetic resonance or mri or mrs or nmr* or angiogra* or tomoangiogra* or imag* or seam* or spectroscop* or tomogra* or elastogra* or examin*) adj3 (abnormal* or lesion* or malformation*).ti,ab. or malformation*).ti,ab. or malformation*.hw.)) use pez or (6 and 9 and (exp case control stud		emczd, emcr
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(lysosomal adj (enzyme or storage) adj (diseases* or disorder*)),ti,ab. axp mitochondrial diseases/ use ppez or exp "disorders of mitochondrial functions"/ use emczd, emcr or ((mitochondrial adj (deficien* or diseases* or disorder*)) or mitochondriopath* or ((electron transport chain or oxidative phosphorylation or respiratory chain) adj2 (deficien* or disease* or disorder*))),ti,ab. amino acid metabolism, inborn errors/ use ppez or "disorders of amino acid and protein metabolism"/ use emczd, emcr or (organic adj (acidemia or aciduria*)),ti,ab. molybdenum cofactor deficiency / use emczd, emcr or (molybdenum adj (co factor or cofactor) adj deficiency),ti,ab. (sulfite oxidase and deficiency),hw. or ((sulfite adj2 oxidase adj2 deficiency) or isod),ti,ab. ((disorder* adj3 (amino acid* or protein*) adj3 metaboli*) or (phenyl ketonuria* or phenylketonuria* or tyrosinemia* or homocystinuria* or non-ketotic hyperglycinemia* or maple syrup urine disease) or (amino acid metablism adj3 inborn error"),ti,ab. (glucose transporter*.sh. and deficien*,hw.) or ((glucose transporter adj3 deficien*) or glut1),ti,ab. (or/10-41) or (abnormal* or lesion* or malformation*),ti,ab. or malformation*.hw. exp epilepsyldi or diagnos*.sh. or (diagnos* or detect* or identif* or indicat* or reveal* or ((epilepsy or seizure) adj protocol*) or yield*),ti,ab. 6 and 9 and 42 and 43 6 and 9 and ((magnetic resonance or mri or mrs or nmr* or angiogra* or tomoangiogra* or imag* or scan* or spectroscop* or formogra* or elastogra* or examin*) adj3 (abnormal* or lesion* or malformation*),ti,ab. 6 and 9 and (exp case control studies/ or exp cohort studies/ or cross-sectional studies/ or epidemiologic studies/ or observational study/ or (case control or (cohort adj (analy* or study or studies)) or cross sectional or (follow up adj (study or studies)) or longitudinal or (observational adj (study or studies)) or retrospective) ti,ab.) and ((abnormal* or lesion* or malformation*),ti,ab. or malformation*,tw.)) use emczd. emczd.emcr.	34	deficien*).ti,ab.
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molybdenum cofactor deficiency / use emczd, emcr or (molybdenum adj (co factor or cofactor) adj deficiency).ti,ab. (sulfite oxidase and deficiency).hw. or ((sulfite adj2 oxidase adj2 deficiency) or isod).ti,ab. ((disorder* adj3 (amino acid* or protein*) adj3 metaboli*) or (phenyl ketonuria* or phenylketonuria* or tyrosinemia* or homocystinuria* or non-ketotic hyperglycinemia* or maple syrup urine disease) or (amino acid metablism adj3 inborn error*)).ti,ab. (glucose transporter*.sh. and deficien*.hw.) or ((glucose transporter adj3 deficien*) or glut1).ti,ab. (or/10-41) or (abnormal* or lesion* or malformation*).ti,ab. or malformation*.hw. exp epilepsy/di or diagnos*.sh. or (diagnos* or detect* or identif* or indicat* or reveal* or ((epilepsy or seizure) adj protocol*) or yield*).ti,ab. 6 and 9 and 42 and 43 6 and 9 and ((magnetic resonance or mri or mrs or nmr* or angiogra* or tomoangiogra* or imag* or scan* or spectroscop* or tomogra* or elastogra* or examin*) adj3 (abnormal* or lesion* or malformation*)).ti,ab. (6 and 9 and (exp case control studies/ or exp cohort studies/ or cross-sectional studies/ or epidemiologic studies/ or observational study/ or (case control or (cohort adj (analy* or study or studies)) or cross sectional or (follow up adj (study or studies)) or longitudinal or (observational adj (study or studies)) or retrospective).ti,ab.) and ((abnormal* or lesion* or malformation*).ti,ab. or malformation*.hw.)) use ppez or (6 and 9 and (exp case control study/ or prospective study/ or retrospective study/ or longitudinal study/ or observational study/ or studies)) or cross sectional or (follow up adj (study or studies)) or longitudinal or (observational adj (study) or studies)) or retrospective).ti,ab.) and ((abnormal* or lesion* or malformation*).ti,ab. or malformation*.hw.)) use emczd, emcr or/44-46 limit 47 to yr="2000 - current" limit 48 to english language ((letter.pt. or letter/ or note.pt. or editorial.pt. or case report/ or case study/ or (letter or comment*).ti.) not (randomized	37	amino acid metabolism, inborn errors/ use ppez or "disorders of amino acid and protein metabolism"/
 ((disorder* adj3 (amino acid* or protein*) adj3 metaboli*) or (phenyl ketonuria* or phenylketonuria* or tyrosinemia* or homocystinuria* or non-ketotic hyperglycinemia* or maple syrup urine disease) or (amino acid metablism adj3 inborn error*)).ti,ab. (glucose transporter*.sh. and deficien*.hw.) or ((glucose transporter adj3 deficien*) or glut1).ti,ab. (or/10-41) or (abnormal* or lesion* or malformation*).ti,ab. or malformation*.hw. exp epilepsy/di or diagnos*.sh. or (diagnos* or detect* or identif* or indicat* or reveal* or ((epilepsy or seizure) adj protocol*) or yield*).ti,ab. 6 and 9 and 42 and 43 6 and 9 and ((magnetic resonance or mri or mrs or nmr* or angiogra* or tomoangiogra* or imag* or scan* or spectroscop* or tomogra* or elastogra* or examin*) adj3 (abnormal* or lesion* or malformation*)).ti,ab. (6 and 9 and (exp case control studies/ or exp cohort studies/ or cross-sectional studies/ or epidemiologic studies/ or observational study/ or (case control or (cohort adj (analy* or study or studies)) or cross sectional or (follow up adj (study or studies)) or longitudinal or (observational adj (study or studies)) or retrospective).ti,ab.) and ((abnormal* or lesion* or malformation*).ti,ab. or malformation*.hw.)) use ppez or (6 and 9 and (exp case control study/ or prospective study/ or retrospective study/ or longitudinal study/ or observational adj (study or studies)) or retrospective study/ or longitudinal study/ or observational adj (study or studies)) or retrospective study/ or (case control or (cohort adj (analy* or study or studies)) or retrospective).ti,ab.) and ((abnormal* or lesion* or malformation*).ti,ab. or malformation*.hw.)) use emczd, emcr or/44-46 limit 47 to yr="2000 - current" limit 48 to english language ((letter.pt. or letter/ or note.pt. or editorial.pt. or case report/ or case study/ or (letter or comment*).ti.) not (randomized controlled trial/ or random*.ti,ab.)) or ((animal/ not human/) or non	38	molybdenum cofactor deficiency / use emczd, emcr or (molybdenum adj (co factor or cofactor) adj
tyrosinemia* or homocystinuria* or non-ketotic hyperglycinemia* or maple syrup urine disease) or (amino acid metablism adj3 inborn error*).ti,ab. 41 (glucose transporter*.sh. and deficien*.hw.) or ((glucose transporter adj3 deficien*) or glut1).ti,ab. 42 (or/10-41) or (abnormal* or lesion* or malformation*).ti,ab. or malformation*.hw. 43 exp epilepsy/di or diagnos*.sh. or (diagnos* or detect* or identif* or indicat* or reveal* or ((epilepsy or seizure) adj protocol*) or yield*).ti,ab. 44 6 and 9 and 42 and 43 45 6 and 9 and ((magnetic resonance or mri or mrs or nmr* or angiogra* or tomoangiogra* or imag* or scan* or spectroscop* or tomogra* or elastogra* or examin*) adj3 (abnormal* or lesion* or malformation*)).ti,ab. 46 (6 and 9 and (exp case control studies/ or exp cohort studies/ or cross-sectional studies/ or epidemiologic studies/ or observational study/ or (case control or (cohort adj (analy* or study or studies)) or cross sectional or (follow up adj (study or studies)) or longitudinal or (observational adj (study or studies)) or retrospective).ti,ab.) and ((abnormal* or lesion* or malformation*).ti,ab. or malformation*.hw.)) use ppez or (6 and 9 and (exp case control study/ or cohort analysis/ or cross-sectional study/ or follow up/ or longitudinal study/ or observational study/ or prospective study/ or retrospective study/ or (case control or (cohort adj (analy* or studies)) or cross sectional or (follow up adj (study or studies)) or longitudinal or (observational adj (study or studies)) or retrospective).ti,ab.) and ((abnormal* or lesion* or malformation*).ti,ab. or malformation*.hw.)) use emczd, emcr 47 or/44-46 48 limit 47 to yr="2000 - current" 49 limit 48 to english language ((letter, pt. or letter/ or note.pt. or editorial.pt. or case report/ or case study/ or (letter or comment*).ti.) not (randomized controlled trial/ or random*.ti,ab.)) or ((animal/ not human/) or nonhuman/ or exp animal experiment/ or exp experimental animal/ or animal model/ or exp rodent/ or (rat or rats or mouse or	39	
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scan* or spectroscop* or tomogra* or elastogra* or examin*) adj3 (abnormal* or lesion* or malformation*)).ti,ab. (6 and 9 and (exp case control studies/ or exp cohort studies/ or cross-sectional studies/ or epidemiologic studies/ or observational study/ or (case control or (cohort adj (analy* or study or studies)) or cross sectional or (follow up adj (study or studies)) or longitudinal or (observational adj (study or studies)) or retrospective).ti,ab.) and ((abnormal* or lesion* or malformation*).ti,ab. or malformation*.hw.)) use ppez or (6 and 9 and (exp case control study/ or cohort analysis/ or cross-sectional study/ or follow up/ or longitudinal study/ or observational study/ or prospective study/ or retrospective study/ or (case control or (cohort adj (analy* or studies)) or cross sectional or (follow up adj (study or studies)) or longitudinal or (observational adj (study or studies)) or retrospective).ti,ab.) and ((abnormal* or lesion* or malformation*).ti,ab. or malformation*.hw.)) use emczd, emcr 47 or/44-46 48 limit 47 to yr="2000 - current" 49 limit 48 to english language 50 ((letter.pt. or letter/ or note.pt. or editorial.pt. or case report/ or case study/ or (letter or comment*).ti.) not (randomized controlled trial/ or random*.ti,ab.)) or ((animal/ not human/) or nonhuman/ or exp animal experiment/ or exp experimental animal/ or animal model/ or exp rodent/ or (rat or rats or mouse or mice).ti.)		
epidemiologic studies/ or observational study/ or (case control or (cohort adj (analy* or study or studies)) or cross sectional or (follow up adj (study or studies)) or longitudinal or (observational adj (study or studies)) or retrospective).ti,ab.) and ((abnormal* or lesion* or malformation*).ti,ab. or malformation*.hw.)) use ppez or (6 and 9 and (exp case control study/ or cohort analysis/ or cross-sectional study/ or follow up/ or longitudinal study/ or observational study/ or prospective study/ or retrospective study/ or (case control or (cohort adj (analy* or study or studies)) or cross sectional or (follow up adj (study or studies)) or longitudinal or (observational adj (study or studies)) or retrospective).ti,ab.) and ((abnormal* or lesion* or malformation*).ti,ab. or malformation*.hw.)) use emczd, emcr 47 or/44-46 48 limit 47 to yr="2000 - current" 49 limit 48 to english language 50 ((letter.pt. or letter/ or note.pt. or editorial.pt. or case report/ or case study/ or (letter or comment*).ti.) not (randomized controlled trial/ or random*.ti,ab.)) or ((animal/ not human/) or nonhuman/ or exp animal experiment/ or exp experimental animal/ or animal model/ or exp rodent/ or (rat or rats or mouse or mice).ti.)	40	scan* or spectroscop* or tomogra* or elastogra* or examin*) adj3 (abnormal* or lesion* or
 limit 47 to yr="2000 - current" limit 48 to english language ((letter.pt. or letter/ or note.pt. or editorial.pt. or case report/ or case study/ or (letter or comment*).ti.) not (randomized controlled trial/ or random*.ti,ab.)) or ((animal/ not human/) or nonhuman/ or exp animal experiment/ or exp experimental animal/ or animal model/ or exp rodent/ or (rat or rats or mouse or mice).ti.) 	46	epidemiologic studies/ or observational study/ or (case control or (cohort adj (analy* or study or studies)) or cross sectional or (follow up adj (study or studies)) or longitudinal or (observational adj (study or studies)) or retrospective).ti,ab.) and ((abnormal* or lesion* or malformation*).ti,ab. or malformation*.hw.)) use ppez or (6 and 9 and (exp case control study/ or cohort analysis/ or cross-sectional study/ or follow up/ or longitudinal study/ or observational study/ or prospective study/ or retrospective study/ or (case control or (cohort adj (analy* or study or studies)) or cross sectional or (follow up adj (study or studies)) or longitudinal or (observational adj (study or studies)) or retrospective).ti,ab.) and ((abnormal* or lesion* or malformation*).ti,ab. or malformation*.hw.)) use emczd, emcr
limit 48 to english language ((letter.pt. or letter/ or note.pt. or editorial.pt. or case report/ or case study/ or (letter or comment*).ti.) not (randomized controlled trial/ or random*.ti,ab.)) or ((animal/ not human/) or nonhuman/ or exp animal experiment/ or exp experimental animal/ or animal model/ or exp rodent/ or (rat or rats or mouse or mice).ti.)		
((letter.pt. or letter/ or note.pt. or editorial.pt. or case report/ or case study/ or (letter or comment*).ti.) not (randomized controlled trial/ or random*.ti,ab.)) or ((animal/ not human/) or nonhuman/ or exp animal experiment/ or exp experimental animal/ or animal model/ or exp rodent/ or (rat or rats or mouse or mice).ti.)		
not (randomized controlled trial/ or random*.ti,ab.)) or ((animal/ not human/) or nonhuman/ or exp animal experiment/ or exp experimental animal/ or animal model/ or exp rodent/ or (rat or rats or mouse or mice).ti.)		
	50	not (randomized controlled trial/ or random*.ti,ab.)) or ((animal/ not human/) or nonhuman/ or exp animal experiment/ or exp experimental animal/ or animal model/ or exp rodent/ or (rat or rats or
	51	50 use emez

#	searches
52	((letter/ or editorial/ or news/ or exp historical article/ or anecdotes as topic/ or comment/ or case report/ or (letter or comment*).ti.) not (randomized controlled trial/ or random*.ti,ab.)) or ((animals not humans).sh. or exp animals, laboratory/ or exp animal experimentation/ or exp models, animal/ or exp rodentia/ or (rat or rats or mouse or mice).ti.)
53	52 use mesz
54	51 or 53
55	49 not 54

Database(s): Cochrane LibraryCochrane Database of Systematic Reviews, Issue 11 of 12, November 2019; Cochrane Central Register of Controlled Trials, Issue 11 of 12, November 2019
Date of last search: 25 November 2019

#	searches						
1	mesh descriptor: [epilepsy] explode all trees						
2	epilep*:ti,ab						
3	(((absence or astatic or atonic or tonic or "tonic clonic") near/2 (seizure* or spasm*)) or (benign near/2 (childhood or neonatal or pediatric or paediatric) near/2 (convulsion* or seizure* or spasm*)) or (benign near/3 convulsion* near/2 centrotemporal near/2 spike*) or ((centralopathic or centrotemporal or "temporal-central" focal) next (convulsion* or seizure*)) or "continous spike wave of slow sleep" or doose* or dravet or ((early or infantile) near/2 myoclonic near/2 encephalopath*) or ((flexor or infantile or neonatal) near/2 (seizure* or spasm*)) or hypsarrhythmia* or "infant* spasm*" or ((jacknife or "jack nife" or lightening or nodding or salaam) next (attack* or convulsion* or seizure* or spasm*)) or (landau near/2 kleffner) or "lennox gastaut" or "massive myoclonia" or (myoclonic near/2 (astatic or atonic)) or (myoclonic near/3 (seizure* or spasm*)) or ((osylvian or postrolandic or roland*) near/2 (convulsion* or seizure* or spasm*)) or "propulsive petit mal" or "spasm in* flexion" or "spasmus nutans" or "west syndrome*"):ti,ab						
4	(bcects or bects or brec or cects or lgs or mae or smei) :ti,ab						
5	{ or #1-#4}						
6	(convulsion* or fits or jerk* or seizure* or spasm*):ti,ab,kw						
7	#5 and #6						
8	mesh descriptor: [magnetic resonance imaging] explode all trees						
9	("magnetic resonance" or mri or mrs or nmr* or ((magnet* or mr or nuclear or nm) near/2 (angiogra* or elastogra* or examin* or imag* or scan* or spectroscop* or tomogra* or tomogragiogra*))):ti,ab						
11	{or #8-#9}						
12	mesh descriptor: [brain injuries] this term only						
13	mesh descriptor: [encephalomalacia] explode all trees						
14	mesh descriptor: [hemorrhage] explode all trees						
15	mesh descriptor: [infarction] this term only						
16	calcification*:kw						
17	mesh descriptor: [vascular malformations] explode all trees						
18	mesh descriptor: [hydrocephalus] explode all trees						
19	mesh descriptor: [edema] explode all trees						
20	mesh descriptor: [brain neoplasms] explode all trees						
21	mesh descriptor: [meningioma] this term only						

#	searches						
22	mesh descriptor: [posterior leukoencephalopathy syndrome] this term only						
23	mesh descriptor: [vasculitis] explode all trees						
24	mesh descriptor: [sinus thrombosis, intracranial] explode all trees						
25	mesh descriptor: [cicatrix] explode all trees						
26	mesh descriptor: [gliosis] this term only						
27	(hippocampus and sclerosis):kw						
28	mesh descriptor: [demyelinating diseases] explode all trees						
29	mesh descriptor: ["malformations of cortical development"] explode all trees						
30	mesh descriptor: [neurocutaneous syndromes] explode all trees						
31	mesh descriptor: [encephalitis] explode all trees						
32	mesh descriptor: [limbic encephalitis] this term only						
33	mesh descriptor: [infection] this term only						
34	mesh descriptor: ["congenital disorders of glycosylation"] this term only						
35	leukodystrophy*:kw.						
36	mesh descriptor: [lysosomal storage diseases] explode all trees						
37	mesh descriptor: [mitochondrial diseases] explode all trees						
38	mesh descriptor: [amino acid metabolism, inborn errors] this term only						
39	(sulfite oxidase and deficiency):kw						
40	("glucose transporter*" and deficien*):kw						
41	((brain* or cerebral) near/2 (abnormal* or damage or lesion* or malformation*)):ti,ab						
42	((brain next (malacia or softening)) or cerebromalacia* or encephalomalacia* or scarring) :ti,ab						
43	(bleeding or (blood next (effusion or loss)) or ha?morrhag* or he?morrhag*):ti,ab						
44	(infarct* or (("thrombo embolic" or thromboembolic) next accident*)):ti,ab						
45	calcification:ti,ab						
46	((vascular next (abnormal* or malformation*)) or ((arteriovenous or "arterio venous") next malformation*) or avm) :ti,ab						
47	("aqueductal stenos?s" or "cerebral ventriculomegal*" or hydrocephal*):ti,ab						
48	(anasarca or dropsy or hydrops or oedema* or edema* or "tissue swelling") :ti,ab						
49	(((brain or cerebral or intracranial or meninges or midline) near/2 (cancer* or metastases or neoplasm* or tumor* or tumour*)) or cerebroma* or mening?oma*):ti,ab						
50	((posterio?r next (leukoencephalopath* or "leuko encephalopath*")) or (posterio?r near/2 reversible encephalopath*) or pres or rpls) :ti,ab						
51	(angiitis or vasculiti*):ti,ab						
52	(!cerebral venous sinus thrombosis! or cvst) :ti,ab						
53	(cicatri?ation or scar* or scarring) :ti,ab						

#	searches							
54	(glios?s or gliomatosis or microgliosis) :ti,ab							
55	((hippocampal or "ammon horn" or hippocampus or incisural or "mesial temporal" or pararhinal) next sclerosis) :ti,ab							
56	ulegyria:ti,ab							
57	(demyelination or (demyelinating near/2 (disorder* or disease*))):ti,ab							
58	((("brain cortext" or cortical) near/2 (dysplasia* or "development malformation*")) or ((abnormal* or malformation*) near/2 "cortical development")) :ti,ab							
59	((neurocutaneous next (disorder* or syndrome*)) or phakoma* or phacomatos*):ti,ab							
60	((allergic next (leukoencephalopath* or "leuko encephalopath*")) or encephaliti* or "limbic encephalit*"):ti,ab							
61	infection*:ti,ab							
62	("carbohydrate deficient glycoprotein syndrome*" or "cdg syndrome*" or ("congenital disorders" near/2 glycosylation) or "glycanosis cdg" or ("carbohydrate deficient" next ("glycoprotein disorders" or "inborn error*"))):ti,ab							
63	((leucodystroph* or "metabolic leucoencephalopa*" or "very long chain") near/3 deficien*):ti,ab							
64	(lysosomal next (enzyme or storage) next (disease* or disorder*)):ti,ab							
65	((mitochondrial next (deficien* or disease* or disorder*)) or mitochondriopath* or (("electron transport chain" or "oxidative phosphorylation" or "respiratory chain") near/2 (deficien* or disease* or disorder*))):ti,ab							
66	(organic next (acidemia or aciduria*)):ti,ab							
67	(molybdenum next ("co factor" or cofactor) next deficiency) :ti,ab							
68	((sulfite near/2 oxidase near/2 deficiency) or isod) :ti,ab							
69	((disorder* near/3 ("amino acid*" or protein*) near/3 metaboli*) or ("phenyl ketonuria*" or phenylketonuria* or tyrosinemia* or homocystinuria* or "non-ketotic hyperglycinemia*" or "maple syrup urine disease") or ("amino acid metabolism" near/3 inborn error*)):ti,ab							
70	(("glucose transporter" near/3 deficien*) or glut1) :ti,ab							
71	(abnormal* or lesion* or malformation*) :ti,ab							
72	malformation*:kw.							
73	{or #12-#72}							
74	MeSH descriptor: [epilepsy] explode all trees and with qualifier(s): [diagnosis - DI]							
75	diagnos*:kw							
76	(diagnos* or detect* or identif* or indicat* or reveal* or ((epilepsy or seizure) next protocol*) or yield*):ti,ab							
77	{or #74-#76}							
78	#7 and #11 and #73 and #77							
79	(("magnetic resonance" or mri or mrs or nmr* or angiogra* or tomoangiogra* or imag* or scan* or spectroscop* or tomogra* or elastogra* or examin*) near/3 (abnormal* or lesion* or malformation*)):ti,ab							
80	#7 and #11 and #79							
81	mesh descriptor: [case control studies] explode all trees							

#	searches
82	mesh descriptor: [cohort studies] explode all trees
83	mesh descriptor: [cross-sectional studies] this term only
84	mesh descriptor: [epidemiologic studies] this term only
85	mesh descriptor: [observational study] this term only
86	("case control" or (cohort next (analy* or study or studies)) or "cross sectional" or ("follow up" next (study or studies)) or longitudinal or (observational next (study or studies)) or retrospective)):ti,ab
87	((abnormal* or lesion* or malformation* or malformation*):ti,ab,kw
88	{or #81-86}
89	#88 and #87
90	#7 and #11 and #89
91	#78 or #80 or #90 with Cochrane Library publication date from Jan 2000 to November 2019

Database(s): DARE; HTA database - CRDDate of last search: 25 November 2019

#	searches
1	mesh descriptor epilepsy explode all trees
2	epilep*
3	(((absence or astatic or atonic or tonic or "tonic clonic") near2 (seizure* or spasm*)) or (benign near2 (childhood or neonatal or pediatric or paediatric) near2 (convulsion* or seizure* or spasm*)) or (benign near3 convulsion* near2 centrotemporal near2 spike*) or ((centralopathic or centrotemporal or "temporal-central" focal) next (convulsion* or seizure*)) or "continous spike wave of slow sleep" or doose* or dravet or ((early or infantile) near2 myoclonic near2 encephalopath*) or ((flexor or infantile or neonatal) near2 (seizure* or spasm*)) or hypsarrhythmia* or "infant* spasm*" or ((jacknife or "jack nife" or lightening or nodding or salaam) next (attack* or convulsion* or seizure* or spasm*)) or (landau near2 kleffner) or "lennox gastaut" or "massive myoclonia" or (myoclonic near2 (astatic or atonic)) or (myoclonic near3 (seizure* or spasm*)) or ((osylvian or postrolandic or roland*) near2 (convulsion* or seizure* or spasm*)) or "propulsive petit mal" or "spasm in* flexion" or "spasmus nutans" or "west syndrome*")
4	(bcects or bects or brec or cects or lgs or mae or smei)
5	#1 or #2 or #3 or #4

Economic

Database(s): MEDLINE & Embase (Multifile) - OVID

Embase Classic+Embase 1947 to 2021 March 31; Ovid MEDLINE(R) and Epub Ahead of Print, In-Process & Other Non-Indexed Citations and Daily 1946 to March 31, 2021 Date of last search: 31 March 2021

Multifile database codes: emczd=Embase Classic+Embase; ppez= MEDLINE(R) and Epub Ahead of Print, In-Process & Other Non-Indexed Citations and Daily

#	searches
1	exp epilepsy/ or exp seizure/ or "seizure, epilepsy and convulsion"/
2	1 use emczd
3	exp epilepsy/ or seizures/ or seizures, febrile/ or exp status epilepticus/

44	and the same of th						
#	searches 3 use ppez						
5	(epilep* or seizure* or convuls*).ti,ab. or (continous spike wave of slow sleep or infant* spasm*).ti,ab.						
6	(seizure and absence).sh. use emczd, emcr or seizures/ use ppez or ((absence adj2 (convulsion* or seizure*)) or ((typical or atypical) adj absenc*) or petit mal* or pyknolepsy or typical absence*).ti,ab.						
7	(atonic seizure or tonic seizure).sh. use emczd, emcr or exp seizures/ use ppez or ((drop or akinetic or atonic or tonic) adj2 (attack* or epileps* or seizure* or convulsion*)).ti,ab. or brief seizure.ti,ab. or (tonic adj3 atonic adj3 (attack* or epileps* or seizure* or convulsion*)).ti,ab.						
8	exp benign childhood epilepsy/ use emczd, emcr or epilepsy, rolandic/ use ppez or (bcects or bects or brec or benign epilepsy or (benign adj2 (childhood or neonatal or pediatric or paediatric) adj2 epileps*) or (benign adj2 (childhood or neonatal or pediatric or paediatric) adj2 (convulsion* or epileps* or seizure* or spasm*)) or (benign adj3 (convulsion* or epileps*) adj2 centrotemporal adj2 spike*) or cects or ((centralopathic or centrotemporal or temporal-central focal) adj (convulsion* or epileps* or seizure*)) or ((osylvian or postrolandic or roland*) adj2 (convulsion* or epileps* or seizure* or spasm*))).ti,ab.						
9	exp generalized epilepsy/ use emczd, emcr or exp epilepsy, generalized/ use ppez						
10	(((akinetic or atonic or central or diffuse or general or generali?ed or idiopathic or tonic) adj3 (epilep* or seizure*)) or ((childhood absence or juvenile absence or myoclonic or myoclonia or myoclonic astatic or myoclonus or gtcs) adj2 epilep*) or (epilepsy adj2 eyelid myoclonia) or (ige adj2 phantom absenc*) or impulsive petit mal or (janz adj3 (epilep* or petit mal)) or jeavons syndrome* or ((janz or lafora or lafora body or lundborg or unverricht) adj2 (disease or syndrome)) or ((jme or jmes) and epilep*) or perioral myoclon*).ti,ab.						
11	infantile spasm/ use emczd, emcr or spasms, infantile/ use ppez or (((early or infantile) adj2 myoclonic adj2 encephalopath*) or ((early or infantile) adj2 epileptic adj2 encephalopath*) or epileptic spasm* or ((flexor or infantile or neonatal) adj2 (seizure* or spasm*)) or generali?ed flexion epileps* or hypsarrhythmia* or ((jacknife or jack nife or lightening or nodding or salaam) adj (attack* or convulsion* or seizure* or spasm*)) or massive myoclonia or minor motor epilepsy or propulsive petit mal or spasm in*1 flexion or spasmus nutans or west syndrome*).ti,ab.						
12	landau kleffner syndrome/ use emczd, emcr, ppez or (dravet or lennox gastaut or lgs or (landau adj2 kleffner) or smei).ti,ab.						
13	lennox gastaut syndrome/ use emczd, emcr or lennox gastaut syndrome/ use ppez or generalized epilepsy/ use emczd, emcr or epileptic syndromes/ use ppez						
14	(child* epileptic encephalopath* or gastaut or lennox or lgs).ti,ab.						
15	myoclonus seizure/ use emczd, emcr or seizures/ use ppez or ((myoclon* adj2 (absence* or epileps* or seizure* or jerk* or progressive familial epilep* or spasm* or convulsion*)) or ((lafora or unverricht) adj2 disease) or muscle jerk).ti,ab.						
16	myoclonic astatic epilepsy/ use emczd, emcr or exp epilepsies, myoclonic/ use ppez or ((myoclonic adj2 (astatic or atonic)) or (myoclonic adj3 (seizure* or spasm*)) or doose* syndrome or mae or generali?ed idiopathic epilepsy).ti,ab. or ((absence or astatic or atonic or tonic or tonic clonic) adj2 (seizure* or spasm*)).ti,ab.						
17	exp epilepsies, partial/ use ppez or exp focal epilepsy/ use emczd, emcr or ((focal or focal onset or local or partial or simple partial) adj3 (epileps* or seizure*)).ti,ab.						
18	severe myoclonic epilepsy in infancy/ use emczd, emcr or exp epilepsies, myoclonic/ use ppez						
19	(dravet*1 or (intractable childhood epilepsy adj2 (generalised tonic clonic or gtc)) or icegtc* or (severe adj2 (myoclonic or polymorphic) adj2 epilepsy adj2 infancy) or smeb or smei).ti,ab.						
20	epilepsy, tonic-clonic/ use ppez or epilepsy, generalized/ use ppez or generalized epilepsy/ use emczd, emcr or grand mal epilepsy/ use emczd, emcr or (((clonic or grand mal or tonic or (tonic adj3 clonic)) adj2 (attack* or contraction* or convuls* or seizure*)) or gtcs or (generali* adj (contraction* or convuls* or insult or seizure*))).ti,ab.						
21	or/2,4-20						
22	exp budgets/ or exp "costs and cost analysis"/ or exp economics, hospital/ or exp economics, medical/ or economics, nursing/ or economics, pharmaceutical/ or economics/ or exp "fees and charges"/ or value of life/						
23	22 use ppez						
24	budget/ or exp economic evaluation/ or exp fee/ or funding/ or health economics/ or exp health care cost/						
25	24 use emczd						
26 27	budget*.ti,ab. cost*.ti.						
28	(economic* or pharmaco economic* or pharmacoeconomic*).ti.						
29	(price* or pricing*).ti,ab.						
30	(cost* adj2 (effective* or utilit* or benefit* or minimi* or unit* or estimat* or variable*)).ab.						
31 32	(financ* or fee or fees).ti,ab. (value adj2 (money or monetary)).ti,ab.						
33	or/23,25-32						

#	searches
34	21 and 33
25	limit 34 to engish language

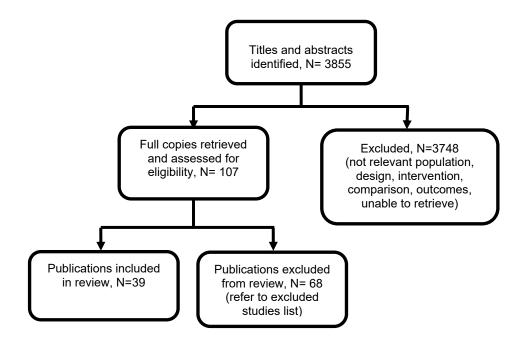
Database(s): NHS Economic Evaluation Database (NHS EED), HTA database – CRD Date of last search: 31 March 2021

	of last search: 31 March 2021						
#	searches						
1	mesh descriptor epilepsy explode all trees						
2	mesh descriptor seizures this term only						
3	mesh descriptor seizures, febrile this term only						
4	mesh descriptor status epilepticus explode all trees						
5	(epilep* or seizure* or convuls*) or ("continous spike wave of slow sleep" or "infant* spasm*")						
6	((absence near2 (convulsion* or seizure*)) or ((typical or atypical) next absenc*) or "petit mal*" or pyknolepsy or "typical absence*")						
7	mesh descriptor seizures explode all trees						
8	((drop or akinetic or atonic or tonic) near2 (attack* or epileps* or seizure* or convulsion*)) or "brief seizure" or (tonic near3 atonic near3 (attack* or epileps* or seizure* or convulsion*))						
9	mesh descriptor epilepsy, rolandic this term only						
10	(bcects or bects or brec or "benign epilepsy" or (benign near2 (childhood or neonatal or pediatric or paediatric) near2 epileps*) or (benign near2 (childhood or neonatal or pediatric or paediatric) near2 (convulsion* or epileps* or seizure* or spasm*)) or (benign near3 (convulsion* or epileps*) near2 centrotemporal near2 spike*) or cects or ((centralopathic or centrotemporal or "temporal-central focal") near (convulsion* or epileps* or seizure*)) or ((osylvian or postrolandic or roland*) near2 (convulsion* or epileps* or seizure* or spasm*)))						
11	mesh descriptor epilepsy, generalized this term only						
12	(((akinetic or atonic or central or diffuse or general or generali?ed or idiopathic or tonic) near3 (epilep* or seizure*)) or (("childhood absence" or "juvenile absence" or myoclonic or myoclonia or "myoclonic astatic" or myoclonus or gtcs) near2 epilep*) or (epilepsy near2 "eyelid myoclonia") or (ige near2 phantom absenc*) or "impulsive petit mal" or (janz near3 (epilep* or "petit mal")) or "jeavons syndrome*" or ((janz or lafora or "lafora body" or lundborg or unverricht) near2 (disease or syndrome)) or ((jme or jmes) and epilep*) or "perioral myoclon*")						
13	mesh descriptor spasms, infantile this term only						
14	(((early or infantile) near2 myoclonic near2 encephalopath*) or ((early or infantile) near2 epileptic near2 encephalopath*) or "epileptic spasm*" or ((flexor or infantile or neonatal) near2 (seizure* or spasm*)) or "generali?ed flexion epileps*" or hypsarrhythmia* or ((jacknife or "jack nife" or lightening or nodding or salaam) next (attack* or convulsion* or seizure* or spasm*)) or "massive myoclonia" or "minor motor epilepsy" or "propulsive petit mal"or "spasm in* flexion" or "spasmus nutans" or "west syndrome*")						
15	mesh descriptor landau kleffner syndrome this term only						
16	(dravet or "lennox gastaut" or lgs or (landau near2 kleffner) or smei)						
17 18	mesh descriptor lennox gastaut syndrome this term only mesh descriptor epileptic syndromes this term only						
19	("child* epileptic encephalopath*" or gastaut or lennox or lgs)						
20	((myoclon* near2 (absence* or epileps* or seizure* or jerk* or "progressive familial epilep*" or spasm* or convulsion*)) or ((lafora or unverricht) near2 disease) or "muscle jerk")						
21 22	mesh descriptor epilepsies, myoclonic explode all trees ((myoclonic near2 (astatic or atonic)) or (myoclonic near3 (seizure* or spasm*)) or "doose* syndrome" or mae or "generali?ed idiopathic epilepsy") or ((absence or astatic or atonic or tonic or "tonic clonic") near2 (seizure* or spasm*))						
23	mesh descriptor epilepsies, partial explode all trees						
24	((focal or "focal onset" or local or partial or "simple partial") near3 (epileps* or seizure*))						
25	mesh descriptor epilepsies, myoclonic this term only						
26	(dravet*1 or ("intractable childhood epilepsy" near2 ("generalised tonic clonic" or gtc)) or icegtc* or (severe near2 (myoclonic or polymorphic) near2 epilepsy near2 infancy) or smeb or smei)						
27	mesh descriptor epilepsy, tonic-clonic this term only						
28	mesh descriptor epilepsy, generalized this term only						
29	(((clonic or "grand mal" or tonic or (tonic near3 clonic)) near2 (attack* or contraction* or convuls* or seizure*)) or gtcs or (generali* next (contraction* or convuls* or insult or seizure*)))						
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						

Appendix C - Clinical evidence study selection

Clinical study selection for: What is the yield of relevant abnormalities detected by MRI in people with epilepsy?

Figure 1: Study selection flow chart



Appendix D – Clinical evidence tables

Clinical evidence tables for review question: What is the yield of relevant abnormalities detected by MRI in people with epilepsy?

Table 4: Clinical evidence tables

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Full citation Alam-Eldeen, M. H., Hasan, N. M. A., Assessment of the diagnostic reliability of brain CT and MRI in pediatric epilepsy patients, Egyptian Journal of Radiology and Nuclear Medicine., 27, 2015 Ref Id 1156238 Country/ies where the study was carried out Egypt Study type Retrospective cohort Aim of the study To assess the role of	Participants Sample size N=181 (74 received CT, 89 received MRI, and 18 received both) Characteristics Age of follow up, years, mean (SD): 4.3 years (range 1 month to 17 years); SD was not reported) Inclusion criteria Not reported Exclusion criteria Those with intracranial tumors and CNS postoperative cases were due to absence of operative and histopathological data	Interventions Interventions MRI 1.5-t	Methods Details Children were clinically diagnosed as having epilepsy and were referred to the Department of Diagnostic Radiology. MR images were reviewed by 2 radiologists for interpretation.	Proportion identified with a clinically relevant abnormality: Vascular: 10/89 Scarring: 3/89 Congenital/ developmental: 33/89 Inflammatory/infective/ immune: 7/89 Proportion identified with a non-epilepsy related abnormality: 8/89	Limitations The quality of this study was assessed using the CEBMA checklist Did the study address a clearly focused question / issue? Yes Is the research method (study design) appropriate for answering the research question? yes Is the method of selection of the subjects (employees, teams, divisions, organizations) clearly described? yes Could the way the sample was obtained introduce (selection)bias? potentially yes as all

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Study dates April 2012 to April 2014 Source of funding Not reported	Participants	Interventions	Methods	Outcomes and Results	Was the sample of subjects representative with regard to the population to which the findings will be referred? unclear Was the sample size based on pre-study considerations of statistical power? no Was a satisfactory response rate achieved? yes Are the measurements (questionnaires) likely to be valid and reliable? yes Was the statistical significance assessed? not applicable
					Are confidence intervals given for the main results? no
					Could there be confounding factors that haven't been accounted for? no
					Can the results be applied to your organization? yes
Full citation	Sample size	Interventions	Details	Results	Limitations

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Ali, A., Akram, F., Khan, G., Hussain, S., Paediatrics Brain Imaging In Epilepsy: Common Presenting Symptoms And Spectrum Of Abnormalities Detected On MRI, Journal of Ayub Medical College, Abbottabad: JAMC, 29, 215-218, 2017 Ref Id 1156894 Country/ies where the study was carried out Pakistan Study type Cross-sectional Aim of the study To assess the yield of MRI abnormalities in people with epilepsy Study dates March 2015 to March 2016 Source of funding Not reported	Characteristics No demographic characteristics were reported Inclusion criteria Those between 28 days and 14 years old with epilepsy Exclusion criteria Not reported	MRI scan 1.5-t	Not reported	Proportion identified with a clinically relevant abnormality: Tumours: 14/209 Vascular: 4/209 Scarring: 3/209 Congenital/ developmental: 16/209 Inflammatory/infective/ immune: 10/209 Metabolic/genetic: 10/209 Proportion identified with a non-epilepsy related abnormality: 8/209	The quality of this study was assessed using the CEBMA checklist Did the study address a clearly focused question / issue? yes Is the research method (study design) appropriate for answering the research question? yes Is the method of selection of the subjects (employees, teams, divisions, organizations) clearly described? no Could the way the sample was obtained introduce (selection)bias? yes Was the sample of subjects representative with regard to the population to which the findings will be referred? unclear Was the sample size based on pre-study considerations of statistical power? no Was a satisfactory response rate achieved? yes

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
					Are the measurements (questionnaires) likely to be valid and reliable? yes Was the statistical significance assessed? not relevant Are confidence intervals given for the main results? no Could there be confounding factors that haven't been accounted for? yes Can the results be applied to your
Full citation Asadi-Pooya, A. A., Sharifzade, M., Lennox-Gastaut syndrome in south Iran: Electro-clinical manifestations, Seizure, 21, 760-763, 2012 Ref Id 1160033 Country/ies where the study was carried out	Sample size N=135 Characteristics Age of follow up, years, mean (SD): 3.2 (3.8) Males, n (%): 83 (61.5) Syndrome type, n (%): Lennox-Gastaut syndrome, 135 (100) Learning disability, n (%): 132 (97) Inclusion criteria	Interventions MRI scan 1.5-t	Details EEG was performed on all patients at the time of referral. No further relevant methods were reported	Results Proportion identified with a clinically relevant abnormality: Congenital/ developmental: 20/135 Metabolic/genetic: 9/135 Proportion identified with a non-epilepsy related abnormality: 1/135	Limitations The quality of this study was assessed using the CEBMA checklist Did the study address a clearly focused question / issue? yes Is the research method (study design) appropriate for answering the research question? yes Is the method of selection of the subjects

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Study type Cross-sectional Aim of the study To assess the prevalence of brain abnormalities in children with Lennox- Gastaut syndrome Study dates September 2008 to May 2012 Source of funding Not reported	Participants Those diagnosed with Lennox-Gastaut syndrome under the care of an epileptologist Exclusion criteria Not reported	Interventions	Methods	Outcomes and Results	Comments (employees, teams, divisions, organizations) clearly described? yes Could the way the sample was obtained introduce (selection)bias? yes Was the sample of subjects representative with regard to the population to which the findings will be referred unclear Was the sample size based on pre-study considerations of statistical power? no Was a satisfactory response rate achieved yes Are the measurements (questionnaires) likely to be valid and reliable? yes Was the statistical significance assessed? not relevant Are confidence intervals given for the main results? no

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
					Could there be confounding factors that haven't been accounted for? yes Can the results be applied to your organization? yes
Full citation Aslan, K., Bozdemir, H., Yapar, Z., Burgut, R., The effect of electrophysiological and neuroimaging findings on the prognosis of juvenile myoclonic epilepsy proband, Neurological Research, 32, 620- 624, 2010 Ref Id 1153393 Country/ies where the study was carried out Turkey Study type Retrospective cohort Aim of the study To report on the clinical, electrophysiological and neuroimaging findings of people with	Sample size N= 32 people with juvenile myoclonic epilepsy Characteristics Age of follow up, years, mean (range): 22 (16 to 37) Males, n (%): 9 (28.12%) Seizure type, n (%): myoclonic + absence + generalised tonic clonic, 22 (68.8); myoclonic + generalised tonic clonic, 8 (25); myoclonic + absence, 2 (6.2) Syndrome type, n (%): 32 (100) juvenile myoclonic epilepsy Response to treatment, n (%): existing diagnosis and controlled, 32 (100) Learning disability, n (%): 3 (9.4) Inclusion criteria Those in whom seizure onset and seizure types were related to juvenile myoclonic epilepsy	Interventions MRI scan 1.5-t	Details People were classified with juvenile myoclonic epilepsy according to ILAE criteria. Diagnosis was based on clinical presentation, history, EEG reports and biochemical analysis. The Porteus Kest was used to evaluate the intelligence quotient. Patients were assessed according to a pre-specified protocol.	Results Proportion identified with a clinically relevant abnormality: Scarring: 1/32 Proportion identified with a non-epilepsy related abnormality: 4/32	Limitations The quality of this study was assessed using the CEBMA checklist Did the study address a clearly focused question / issue? yes Is the research method (study design) appropriate for answering the research question? yes Is the method of selection of the subjects (employees, teams, divisions, organizations) clearly described? no Could the way the sample was obtained introduce (selection)bias? unclear as the way the sample was obtained was not reported

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Study details invenile myoclonic epilepsy Study dates Not reported Source of funding Not reported	Participants Those taking entiepileptic medication >1 year Those without CNS developmental abnormality (with or without progressive learning disability) Exclusion criteria Not rerported	Interventions	Methods	Outcomes and Results	Was the sample of subjects representative with regard to the population to which the findings will be referred? unlcear Was the sample size based on pre-study considerations of statistical power? no Was a satisfactory response rate achieved? yes Are the measurements (questionnaires) likely to be valid and reliable? yes Was the statistical significance assessed? not relevant Are confidence intervals given for the main results? no Could there be confounding factors that haven't been accounted.
					confounding factors tha

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Full citation Bakhsh, A., Value of neuroimaging in epilepsy: An experience from Pakistan, Journal of Neurosciences in Rural Practice, 4, S35-S39, 2013 Ref Id 1153420 Country/ies where the study was carried out Pakistan Study type Prospective cohort Aim of the study To evaluate structural brain lesions in patients with epilepsy Study dates Not reported Source of funding Not reported	Sample size N=366, n=339 received CT scans and n=44 received MRI scans Characteristics Age of follow up, years, mean (SD): 19.5 (SD not reported) Males, n (%): 240 (65.5) Seizure type, n (%): generalised tonic clonic, n=282 (77.04); complex partial seizure leading to generalised tonic clonic, n=70 (19.12); partial motor fits leading to generalised tonic clonic, n=10 (2.7); juvenile myoclonic epilepsy, n=2 (0.5); complex partial seizures, n=2 (0.5) Learning disability, n (%): 19 (5.1) Inclusion criteria Patients with epilepsy, regardless of cause, type or neurological status Exclusion criteria Those <1 year old Those with a first seizure, pseudoseizures, pregnancy, seizures secondary to any metabolic disorders, seizures with a frequency of only 1 per annum	Interventions Interventions MRI scan 1-t	Details Diagnosis of epilepsy was made based on clinical history. MRI scans were done without contrast due to budget constraints. No protocols of hipocampus volumetry was done in any MRI scans. Scans were interpreted by general radiologists	Results Proportion identified with a clinically relevant abnormality: Tumours: 3/44 Vascular: 4/44 Scarring: 9/44 Proportion identified with a non-epilepsy related abnormality: 3/44	Limitations The quality of this study was assessed using the CEBMA checklist Did the study address a clearly focused question / issue? yes Is the research method (study design) appropriate for answering the research question? yes Is the method of selection of the subjects (employees, teams, divisions, organizations) clearly described? no Could the way the sample was obtained introduce (selection)bias? yes Was the sample of subjects representative with regard to the population to which the findings will be referred? unclear Was the sample size based on pre-study considerations of statistical power? no

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
					Was a satisfactory response rate achieved? yes
					Are the measurements (questionnaires) likely to be valid and reliable? yes
					Was the statistical significance assessed? not relevant
					Are confidence intervals given for the main results? no
					Could there be confounding factors that haven't been accounted for? yes
					Can the results be applied to your organization? yes
Full citation Benson, J. C., Chiu, S., Flemming, K., Nasr, D. M., Lanzino, G., Brinjikji, W., MR	Sample size N=57 with a seizure at initial clinical presentation Characteristics	Interventions MRI scans 1.5-t and 3-t	Details Two blinded reviewers assessed the patients's	Results Proportion identified with a clinically relevant abnormality: Vascular: 57/57	Limitations The quality of this study was assessed using the CEBMA checklist
characteristics of unruptured intracranial arteriovenous	Age of follow up, years, mean (SD): 35.9 (SD not reported) Males, n (%): 30 (52.6)		characteristics, including imaging, lesion locality, and	Scarring: 38/57 Proportion identified with a non-epilepsy related	Did the study address a clearly focused question / issue? yes
malformations associated with seizure as initial	Syndrome type, n (%): 57 (100) arteriovenous malformation with 1		characteristics of AVMs.	abnormality: 12/57	Is the research method (study design) appropriate for
clinical presentation, Journal of	seizure at first clinical presentaition		People were assessed		answering the research question? yes

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Ref Id 1157597 Country/ies where the study was carried out US Study type Retrospective cohort Aim of the study To assess MRI characteristics in people with intracranial arteriovenous malformations associated with seizures at initial clinical presentation Study dates 1 January 2000 to 31 December 2016 Source of funding No specific source of funding was reported	Inclusion criteria Those diagnosed with AVM at the study's institution within the provided timeframe Those with peri-AVM on T2 imaging were also included provided they had no previous history of AVM and they had never had any imaging evidence of acure or subacute haemorrhage Exclusion criteria Those with extracranial AVM Those with AVM with history of acute rupture People who had undergone treatment for AVM AVM not identified on MRI		according to a pre-specified protocol, although 25 different scanners were used within the institution.		Is the method of selection of the subjects (employees, teams, divisions, organizations) clearly described? yes Could the way the sample was obtained introduce (selection)bias? yes Was the sample of subjects representative with regard to the population to which the findings will be referred? unclear Was the sample size based on pre-study considerations of statistical power? no Was a satisfactory response rate achieved yes Are the measurements (questionnaires) likely to be valid and reliable? yes Was the statistical significance assessed? not relevant

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
					Are confidence intervals given for the main results? no Could there be confounding factors that haven't been accounted for? yes (different MRI
					scans with different strenght of magnet were used)
					Can the results be applied to your organization? yes
					Other information Note: presence of AVM part of the inclusion criteria, which may overstimate the yield of vascular abnormalities
Full citation Berg, A. T., Testa, F.	Sample size N= 388 children with newly	Interventions MRI, strength of	Details Children were	Results Proportion identified with a	Limitations The quality of this study
M., Levy, S. R.,	diagnosed epilepsy	magnet was not	entered in the	clinically relevant	was assessed using
Shinnar, S., Neuroimaging in	Characteristics	reported	study when they were first	<u>abnormality:</u> Tumours: 2/388	the CEBMA checklist
children with newly diagnosed epilepsy: A community-based study, Pediatrics, 106,	Age at seizure onset, years, median: 5.7 (range/IQR was not reported)		diagnosed with epilepsy. Etiology was based on medical records	Vascular: 11/388 Scarring: 5/388 Congenital/ developmental: 41/388	Did the study address a clearly focused question / issue? yes
527-532, 2000	Inclusion criteria		and information obtained from	Inflammatory/infective/ immune: 3/388	Is the research method (study design)
Ref Id 1153473	Those between 1 month and 15 years		parents. MRI was considered if it	Metabolic/genetic: 15/388 Proportion identified with a non-epilepsy related	appropriate for answering the research question? yes
	Exclusion criteria		was ordered as	abnormality: 15/388	

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Country/ies where the study was carried out US Study type Retrospective cohort Aim of the study To assess the yield of neuroimaging in people with epilepsy Study dates 1993 to 1997 Source of funding National Institutes of Health	• Not reported	Interventions	Methods part of the initial assessment or if these have been done before the onset of epilepsy.	Outcomes and Results	Is the method of selection of the subjects (employees, teams, divisions, organizations) clearly described? unclear - resons for inclusion/ exclusion are not provided in detail Could the way the sample was obtained introduce (selection)bias? yes Was the sample of subjects representative with regard to the population to which the findings will be referred? unclear Was the sample size based on pre-study considerations of statistical power? no Was a satisfactory response rate achieved? yes Are the measurements (questionnaires) likely to be valid and reliable? yes Was the statistical significance assessed?

Are confidence intervals given for the main results? no Could there be confounding factors that haven't been accounted for?' yes Esting, L. E., Mory, S. B., Lopes-Cendes, I., I., L. M., Guerreiro, C. A. M., Gendes, F., MRI reveals structural abnormalities in patients with dicipathic generalized epilepsy, Neurology, 67, 848-852, 2006 Ref Id 1158776 Country/ies where the study was carried out Fiszarial Properties of the study was carried out Fiszarial Study type Prospective cohort study Study type Prospective cohort study Inclusion Criteria Interventions MRI scan 2.0-t. MRI protocol was used in all patients. Details Proportion identified with a clinical history of generalized epilepsy, Neurology, 67, 848-852, 2006 Ref Id 1158776 Study type Prospective cohort study Inclusion criteria Interventions MRI scan 2.0-t. MRI protocol was used in all patients. Proportion identified with a clinical history of generalized epilepsy, 134 (100) Service of the wash of the main results? No Could there be confounding factors that haven't been accounted for? yes Can the results be applied to your organizations? Details Proportion identified with a clinical history of generalized epilepsy. To comparize for an incomplete proportion identified with a non-epilepsy related abnormality. 2/134 Study type Prospective cohort study Interventions MRI scan 2.0-t. MRI protocol was used in all patients. Sample size in the results be applied to your organization? Yes Details A pre-specified MRI protocol was used in all patients. A pre-specified MRI protocol was used in all patients. A pre-specified MRI protocol was used in all patients. Scarried out protocol was used in all patients. Subject on the study with a clinical history of generalized epilepsy, 11 (52.9); and the protocol was assessed using the CEBMA checklist organizations. Did the study design organizations. Did the study design organizations. Did the study of this study in the protocol was used in all patients. Subject of the subject organizations organizations	Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Betting, L. E., Mory, S. B., Lopes-Cendes, I., Li, L. M., Guerreiro, M. M., Guerreiro, C. A. M., Cendes, F., MRI reveals structural abnormalities in patients with idiopathic generalized epilepsy, Neurology, 67, 848-852, 2006 Ref Id 1158776 Country/les where the study was carried out Brazil Study type Prospective cohort study N=134 N=134 MRI scan 2.0-t MRI scan 2.0-t MRI scan 2.0-t A pre-specified MRI protocol was used in all patients. Scarring: 11/134 Congenital/developmental: 3/134 Congenital/developmental: 3/134 Seizure type n (%): 51 (38.05) Seizure type n (%): idiopathic generalised epilepsy, 134 (100) Syndrome type, n (%): juvenile myoclonic epilepsy, 22 (16.4); generalised tonic clonic, 41 (30.5) Study type Prospective cohort study MRI scan 2.0-t A pre-specified MRI protocol was used in all patients. Scarring: 11/134 Congenital/developmental: 3/134 Seizure type n (%): idiopathic generalised epilepsy, 134 (100) Syndrome type, n (%): juvenile myoclonic epilepsy, 71 (52.9); absence epilepsy, 72 (16.4); generalised tonic clonic, 41 (30.5) Study type Prospective cohort study Proportion identified with a clinically relevant abnormality: Scarring: 11/134 Congenital/developmental: 3/134 Seizure type n (%): idiopathic generalised epilepsy, 134 (100) Syndrome type, n (%): juvenile myoclonic epilepsy, 71 (52.9); absence epilepsy, 72 (16.4); generalised tonic clonic, 41 (30.5) Study type Prospective cohort study Proportion identified with a clinically relevant abnormality: Scarring: 11/134 Congenital/developmental: 3/134 Seizure type n (%): idiopathic generalised abnormality: Scarring: 11/134 Congenital/developmental: 3/134 Seizure type n (%): identified with a non-epilepsy related abnormality: Scarring: 11/134 Congenital/developmental: 3/134 Seizure type n (%): identified with a non-epilepsy related abnormality: Scarring: 11/134 Congenital/developmental: 3/134 Step reverse type n (%): identified with a cliently abnormality: Scarring: 11/134 Seizure type n (%): identified with a						given for the main results? no Could there be confounding factors that haven't been accounted for? yes Can the results be applied to your
Aim of the study Exclusion criteria	Betting, L. E., Mory, S. B., Lopes-Cendes, I., Li, L. M., Guerreiro, M. M., Guerreiro, C. A. M., Cendes, F., MRI reveals structural abnormalities in patients with idiopathic generalized epilepsy, Neurology, 67, 848-852, 2006 Ref Id 1158776 Country/ies where the study was carried out Brazil Study type Prospective cohort	Characteristics Age at seizure onset, years, mean (SD): 28 (9) Age of follow up, years, mean (SD): 13 (7) Males, n (%): 51 (38.05) Seizure type n (%): idiopathic generalised epilepsy, 134 (100) Syndrome type, n (%): juvenile myoclonic epilepsy, 71 (52.9); absence epilepsy, 22 (16.4); generalised tonic clonic, 41 (30.5) Inclusion criteria Those with a clinical history of generalised seizures		A pre-specified MRI protocol was used in all	Proportion identified with a clinically relevant abnormality: Scarring: 11/134 Congenital/developmental: 3/134 Proportion identified with a non-epilepsy related	The quality of this study was assessed using the CEBMA checklist Did the study address a clearly focused question / issue? yes Is the research method (study design) appropriate for answering the research question? yes Is the method of selection of the subjects (employees, teams, divisions, organizations) clearly described? yes Could the way the sample was obtained introduce

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
To assess MRI findings in people with idiopathic generalised epilepsy Study dates 2000 to 2005 Source of funding Amparo a Pesquisa do Estado de Sao Paulo (FAPESP) and Coordenacao de Aperfeicoamento de Pessoal de Nivel Superior (CAPES)	Those above 50 years old Those with suspected focal seizure				Was the sample of subjects representative with regard to the population to which the findings will be referred? unclear Was the sample size based on pre-study considerations of statistical power? no Was a satisfactory response rate achieved? yes Are the measurements (questionnaires) likely to be valid and reliable? yes Was the statistical significance assessed? not relevant Are confidence intervals given for the main results? no Could there be confounding factors that haven't been accounted for? yes Can the results be applied to your organization? yes
Full citation	Sample size	Interventions	Details	Results	Limitations

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Bruno, V., Klein, J. P.,	N=217	MRI scan 1.5-t	People were	Proportion identified with a	The quality of this study
Nidup, D., Nirola, D.			recruited from an	clinically relevant	was assessed using
K., Tshering, L., Deki,	Characteristics		existing epilepsy	abnormality:	the CEBMA checklist
S., Clark, S. J., Linn,	Age of follow up, years, mean		registry and	Tumours	Did the study address a
K. A., Shinohara, R.	(SD):		referred through	Children: 0/54	clearly focused question
T., Dorji, C., Pokhrel,	Children: 11.7 (8)		healthcare	Adults: 4/163	/ issue? yes
D. R., Dema, U.,	Adults: 30.2 (11)		profesionals. A	Overall: 4/217	
Mateen, F. J., Yield of			neurologist or		Is the research method
Brain MRI in Clinically	Males, n (%):		psyshiatrist	Vascular	(study design)
Diagnosed Epilepsy in	Children: 26 (48.14)		evaluated each	Children: 6/54	appropriate for
the Kingdom of	Adults: 67 (41.10)		participant and	Adults: 9/163	answering the research
Bhutan: A Prospective	Decrease to the observation (0/):		confirmed the	Overall: 13/217	question? yes
Study, Annals of	Response to treatment, n (%):		clinical diagnosis.	Caamina	le the weetherd of
Global Health, 83, 415-422, 2017	217 (100) existing diagnosis and resistant to treatment			Scarring Children: 0/54	Is the method of selection of the subjects
413-422, 2017	resistant to treatment			Adults: 2/163	(employees, teams,
Ref Id	Inclusion criteria			Overall: 2/217	divisions, organizations)
1156928	Bhutan residents			Overall. 2/217	clearly described? yes
1100020				Congenital/	crearry accompany yes
Country/ies where	 Diagnosis of epilepsy 			developmental	Could the way the
the study was	Production office			Children: 14/54	sample was obtained
carried out	Exclusion criteria			Adults: 15/163	introduce
Bhutan	• Those with non-epileptic epilepsy events			Overall: 29/217	(selection)bias? yes
Study type	 Those with febrile seizures 			Inflammatory/infective/	Was the sample of
Prospective cohort	Those with alcohol or metabolic-			immune	subjects representative
	related seizures			Children: 1/54	with regard to the
Aim of the study	Those under 5 not needing an			Adults: 25/163	population to which the
To assess the yield of brain MRI in people	MRI for clinical reasons			Overall: 26/217	findings will be referred? unclear
with epilepsy				Metabolic/genetic	
				Children: 0/54	Was the sample size
Study dates				Adults: 1/163	based on pre-study
July 2014 to				Overall: 1/217	considerations of
December 2015					statistical power? no
				Proportion identified with a	
Source of funding				non-epilepsy related	Was a satisfactory
Government of				abnormality:	response rate achieved?
Canada; Thrasher				Children: 5/54	yes

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Foundation; Charles Hood Foundation. Two authors were partially funded by a grant				Adults: 23/163 Overall: 28/217	Are the measurements (questionnaires) likely to be valid and reliable? yes
					Was the statistical significance assessed? not relevant
					Are confidence intervals given for the main results? no
					Could there be confounding factors that haven't been accounted for? yes
					Can the results be applied to your organization? yes
					Other information Note: neurocysticercosis is endemic to Bhutan, the infections detected in MRI were all neurocysticercosis, which may overestimate the yield of MRI for infections in this group
Full citation Byars, A. W., deGrauw, T. J., Johnson, C. S., Fastenau, P. S.,	Sample size N= 249 Characteristics	Interventions MRI scans. Strenght magnet varied	Details Participants had their MRI within 6 months of the first	Results Proportion identified with a clinically relevant abnormality: Scarring: 29/249	Limitations The quality of this study was assessed using the CEBMA checklist

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Perkins, S. M., Egelhoff, J. C., Kalnin, A., Dunn, D. W., Austin, J. K., The	Age of follow up, years, mean (SD): 9.6 (2.5) Males, n (%): 198 (79.5)	between 0.5 and 1.5-t	seizure (median 1.3 months).	Congenital/developmental: 6/249 Proportion identified with a	Did the study address a clearly focused question / issue? yes
association of MRI findings and neuropsychological functioning after the first recognized seizure, Epilepsia, 48, 1067-74, 2007	Seizure type: mixed Syndrome type: mixed Inclusion criteria		neuroradiologists to EEG findings reviewed the data. Scanners were done according to a standardised	non-epilepsy related abnormality: 5/249	Is the research method (study design) appropriate for answering the research question? yes
Ref Id 1158973	 Those aged 6 to 14 years old Those with a first recognised seizure within the past 3 months 		seizure protocol.		selection of the subjects (employees, teams, divisions, organizations) clearly described? yes
Country/ies where the study was carried out US Study type	 Exclusion criteria Those whose seizure provoked from an acute situational etiology such as infection, toxin, trauma 				Could the way the sample was obtained introduce (selection)bias? yes
Prospective cohort Aim of the study To assess the prevalence of MRI abnormalities in people with epilepsy	or a mass lesion Those with chronic co-occurring conditions				Was the sample of subjects representative with regard to the population to which the findings will be referred? unclear
after their first seizure Study dates July 2000 to June 2004					Was the sample size based on pre-study considerations of statistical power? no
Source of funding National Institute of Neurological					Was a satisfactory response rate achieved? yes
Disorders and Stroke					Are the measurements (questionnaires) likely to

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Study details	Participants	Interventions	Methods	Outcomes and Results	be valid and reliable? yes Was the statistical significance assessed? no Are confidence intervals given for the main results? no Could there be confounding factors that haven't been accounted for? yes Can the results be applied to your organization? yes Other information Scans were done within 3 months from the onset of the first seizure, therefore the age at follow-up and onset
Full citation Coryell, J., Gaillard,	Sample size N=714 infants with early life	Interventions MRI scan 1.5 or	Details For each of the	Results Proportion identified with a	were very close in time Limitations The quality of this study
W. D., Shellhaas, R. A., Grinspan, Z. M., Wirrell, E. C., Knupp, K. G., Wusthoff, C. J.,	epilepsy Characteristics Age at seizure onset, months,	3-t, results have not been reported separately	participating centres, paediatric epileptologists,	clinically relevant abnormality: Vascular: 55/ 714 Scarring: 9/714	was assessed using the CEBMA checklist Did the study address a
Keator, C., Sullivan, J. E., Loddenkemper, T., Patel, A., Chu, C. J., Massey, S., Novotny,	mean (SD): 11.1 (SD not reported) Age of follow up, months, mean (SD): 12.7 (SD not reported)		identified the children relevant for inclusion.	Congenital/ developmental: 109/714 Inflammatory/infective/ immune: 8/714	clearly focused question / issue? yes Is the research method
E. J., Saneto, R. P.,	(Metabolic/genetic: 5/714	(study design)

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Berg, A. T., Neuroimaging of early life epilepsy, Pediatrics, 142 (3) (no pagination), 2018 Ref Id 1098077 Country/ies where the study was carried out US Study type Prospective cohort Aim of the study To assess the yield of MRI abnormalities in infant with early life epilepsy Study dates 2012-2015 Source of funding Pediatric Epilepsy Research Foundation in Dallas, Texas.	Inclusion criteria Infants with a first seizure before their 3rd birthday and with a diagnosis of epilepsy established before 42 months of age Exclusion criteria Not reported		Researchers obtained relevant data from medical records. Scans done within 1 year of first seizure, were reviewed by a lead study coordinator and the principal study investigator.	Proportion identified with a non-epilepsy related abnormality: 20/714	appropriate for answering the research question? yes Is the method of selection of the subjects (employees, teams, divisions, organizations) clearly described? yes Could the way the sample was obtained introduce (selection)bias? yes Was the sample of subjects representative with regard to the population to which the findings will be referred? unclear as subjects were referred from tertiary centers and this may overestimate the severity of some cases Was the sample size based on pre-study considerations of statistical power? No Was a satisfactory response rate achieved? yes Are the measurements (questionnaires) likely to be valid and reliable? yes

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
					Was the statistical significance assessed? not relevant Are confidence intervals given for the main results? no Could there be confounding factors that haven't been accounted for? yes Can the results be applied to your organization? yes
Full citation Craven, I., Griffiths, P. D., Bhattacharyya, D., Grunewald, R. A., Hodgson, T., Connolly, D. J. A., Coley, S. C., Batty, R., Romanowski, C. A. J., Hoggard, N., 3.0 T MRI of 2000 consecutive patients with localisation- related epilepsy, British Journal of Radiology, 85, 1236- 1242, 2012 Ref Id 1160064	Sample size N=2000 Characteristics Age of follow up, years, median (range): 23 (25 to 48 years) Males, n (%): 922 (46.1) Inclusion criteria Not reported Exclusion criteria Those with generalised epilepsy and those with first seizures	Interventions MRI scan 3.0-t	Details Patients were referred to the neuroscience facility from a catchment area of 2.3 million people. People were scanned with a protocol only used for people with epilepsy. Examinations were reviewed by experienced neuroradiologists and whether findings were related or not to	Results Proportion identified with a clinically relevant abnormality: Tumours: 20/2000 Vascular: 33/2000 Scarring: 248/2000 Congenital/ developmental: 73/2000 Inflammatory/infective/ immune: 4/2000 Metabolic/genetic: 6/2000 Proportion identified with a non-epilepsy related abnormality: 326/2000	Limitations The quality of this study was assessed using the CEBMA checklist Did the study address a clearly focused question / issue? yes Is the research method (study design) appropriate for answering the research question? yes Is the method of selection of the subjects (employees, teams, divisions, organizations) clearly described? yes

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Country/ies where the study was carried out UK			epilepsy, was discussed in a "multidisciplinaty epilepsy meeting"		Could the way the sample was obtained introduce (selection)bias? no
Study type Retrospective cohort Aim of the study To evaluate the yield of radiological abnormalities in					Was the sample of subjects representative with regard to the population to which the findings will be referred? yes
people with localised seizures Study dates					Was the sample size based on pre-study considerations of statistical power? no
January 2005 to February 2011 Source of funding					Was a satisfactory response rate achieved? yes
Not reported					Are the measurements (questionnaires) likely to be valid and reliable? yes
					Was the statistical significance assessed? not relevant
					Are confidence intervals given for the main results? no
					Could there be confounding factors that haven't been accounted for? no

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
					Can the results be applied to your organization? yes
Full citation Das, P., Bindu, P. S., Bharath, R. D., Saini, J. S., Prasad, C., Sinha, S., MRI observations in children with epilepsy: Experience from a large cohort, Journal of Pediatric Epilepsy, 2, 223-228, 2013 Ref Id 1153713 Country/ies where the study was carried out India Study type Cross-sectional Aim of the study To assess the yield of MRI abnormalities in people with epilepsy Study dates August 2009 to January 2011 Source of funding	Sample size N=144 Characteristics Age at seizure onset, years, mean (SD): 2.91 (3.30 years) Age of follow up, years, mean (SD): 5.87 (4.19 years) Males, n (%): 73 (50.69) Seizure type, n (%): partial in n=67 (46.5); generalised in n=72 (50); and unclassified in n=5 (3.4) Syndrome type n (%): structural/metabolic (symptomatic), n=95 (65.9); unknown (cryptogenic), n= 45 (31.25); genetic (idiopathic), n=6 (4.1) Learning disability, n (%): 71 (49.3) Inclusion criteria Not reported Exclusion criteria Those with neonatal or febrile seizures	Interventions MRI scan 1.5 or 3-t	Details The study was conducted in the departments of nerorology and neuroradiology in a teaching hospital. Seizure type was classified according to ILAE criteria/ revised classification of epilepsy and epilepsy syndromes. Patients underwent EEG and MRI according to a standardised protocol.	Results Proportion identified with a clinically relevant abnormality: Tumours: 4/144 Vascular: 10/144 Scarring: 17/144 Congenital/ developmental: 20/144 Inflammatory/infective/imm une: 5/144 Metabolic/genetic: 1/144 Proportion identified with a non-epilepsy related abnormality: 29/144	Limitations The quality of this study was assessed using the CEBMA checklist Did the study address a clearly focused question / issue? yes Is the research method (study design) appropriate for answering the research question? yes Is the method of selection of the subjects (employees, teams, divisions, organizations) clearly described? yes Could the way the sample was obtained introduce (selection)bias? yes Was the sample of subjects representative with regard to the population to which the findings will be referred? unclear

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Not reported					Was the sample size based on pre-study considerations of statistical power? no
					Was a satisfactory response rate achieved? yes
					Are the measurements (questionnaires) likely to be valid and reliable? yes
					Was the statistical significance assessed? not relevant
					Are confidence intervals given for the main results? no
					Could there be confounding factors that haven't been accounted for? yes
					Can the results be applied to your organization? yes
Full citation Dirik, M. A., Sanlidag, B., Magnetic resonance imaging	Sample size N=222 Characteristics	Interventions MRI scan 1.5-t or 3-t	Details Children were recruited from the department of	Proportion identified with a clinically relevant abnormality:	Limitations The quality of this study was assessed using the CEBMA checklist
and interictal electroencephalograp hy findings in newly diagnosed epileptic children, Journal of	Age at seizure onset, months, mean (SD): 48 (SD not reported) Males, n (%): 147 (66.2)		paediatric neurology. MRI protocol was standardised	Tumours: 1/222 Vascular: 3/222 Scarring: 23/222 Congenital/developmental: 25/222	Did the study address a clearly focused question / issue? yes

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Study details Clinical Medicine, 7 (6) (no pagination), 2018 Ref Id 1157305 Country/ies where the study was carried out Cyprus Study type Retrospective cohort Aim of the study To assess the prevalence of MRI lesions in children with newly diagnosed epilepsy Study dates Not reported Source of funding Not reported	Inclusion criteria Those aged between 3 months and 18 years of age Exclusion criteria Not reported	Interventions	Methods	Proportion identified with a non-epilepsy related abnormality: 9/222	Is the research method (study design) appropriate for answering the research question? yes Is the method of selection of the subjects (employees, teams, divisions, organizations) clearly described? yes Could the way the sample was obtained introduce (selection)bias? yes Was the sample of subjects representative with regard to the population to which the findings will be referred? yes Was the sample size based on pre-study considerations of statistical power? no Was a satisfactory response rate achieved? yes Are the measurements (questionnaires) likely to be valid and reliable?

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
					Was the statistical significance assessed? no
					Are confidence intervals given for the main results? no
					Could there be confounding factors that haven't been accounted for? yes
					Can the results be applied to your organization? yes
Full citation Dura-Trave, T., Yoldi- Petri, M. E., Esparza- Estaun, J., Gallinas- Victoriano, F., Aguilera-Albesa, S., Sagastibelza- Zabaleta, A., Magnetic resonance imaging abnormalities in children with epilepsy, European Journal of Neurology, 19, 1053-1059, 2012 Ref Id 1160077	Sample size N=457 Characteristics Age, years, at the time of diagnosis: 1 month to 15 years old Males. n (%): 233 (51) Syndrome type: mixed (West Syndrome, myoclonic epilepsy in infancy, Dravet syndrome) Inclusion criteria Those between 1 month and 15 years of age at the time of diagnosis	Interventions MRI scan (strength of magnet was not reported)	Details Medical records from children referred to the neuropaediatric department of reference within the region where the study was conducted were included. Children were scanned according to a standardised protocol	Results Proportion identified with a clinically relevant abnormality: Tumours: 2/457 Vascular: 12/457 Scarring: 76/457 Congenital/developmental: 33/457 Proportion identified with a non-epilepsy related abnormality: 47/457	Limitations The quality of this study was assessed using the CEBMA checklist Did the study address a clearly focused question / issue? yes Is the research method (study design) appropriate for answering the research question? yes Is the method of selection of the subjects (employees, teams,
Country/ies where the study was carried out Spain	Exclusion criteria				divisions, organizations) clearly described? yes

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Study type Retrospective cohort Aim of the study To assess the proportion of MRI abnormalities in children with epilepsy Study dates Not reported Source of funding No specific grant was received	Participants Those with neonatal seizures only, febrile seizures, and other acute symptomatic seizures Participants Those with neonatal seizures only, febrile seizures, and other acute symptomatic seizures	Interventions	Methods	Outcomes and Results	Comments Could the way the sample was obtained introduce (selection) bias? yes Was the sample of subjects representative with regard to the population to which the findings will be referred? yes Was the sample size based on pre-study considerations of statistical power? No Was a satisfactory response rate achieved? yes Are the measurements (questionnaires) likely to be valid and reliable? yes Was the statistical significance assessed? not relevant Are confidence intervals given for the main results? no Could there be confounding factors that haven't been accounted for? yes

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
					Can the results be applied to your organization? yes
Full citation Ekici, F., Tekbas, G., Onder, H., Gumus, H., Cetincakmak, M. G., Balik, S. K., Acar, A., Hamidi, C., Bilici, A., Comparison of 3.0-T MRI findings in drug resistant and non-resistant adult epileptic patients, Neurology Psychiatry and Brain Research, 19, 42-47, 2013 Ref Id 1155672 Country/ies where the study was carried out Turkey Study type Retrospective cohort Aim of the study To assess the prevalence of MRI abnormalities in a sample of people with epilepsy Study dates December 2009 - October 2011	Characteristics Age of follow up: range 18 to 82; mean 31.3 Males, n (%): 150 (56.8) Response to treatment, n (%): existing diagnosis and resistant to medical treatment, n=94 (35); existing diagnosis (non-resistant to medical treatment), n= 170 (64.3%) (unclear if patients had an existing diagnosis) Inclusion criteria Not reported Exclusion criteria Not reported	Interventions MRI scan 3-t	Details Diagnosis was established based on the clinical and EEG findings by one neurologist. Those who received a single antiepileptic drug to control seizures were considered non-resistant to treatment and those who had 2 or more seizures per month for a period of more than 2 years with 2 or more antiepileptic drugs attending the intractable epilepsy outpatient clinic. All patients underwent MRI sequences according to a standardised protocol.	Resistant to medical treatment Proportion identified with a clinically relevant abnormality: Tumours: 4/94 Vascular: 7/94 Scarring: 39/94 Congenital/developmental: 10/94 Proportion identified with a non-epilepsy related abnormality: 0/94 Non-resistant to medical treatment Proportion identified with a clinically relevant abnormality: Tumours: 0/170 Vascular: 3/170 Scarring: 35/170 Congenital/developmental: 0/170 Proportion identified with a non-epilepsy related abnormality:4/170	Limitations The quality of this study was assessed using the CEBMA checklist Did the study address a clearly focused question / issue? yes Is the research method (study design) appropriate for answering the research question? yes Is the method of selection of the subjects (employees, teams, divisions, organizations) clearly described? yes Could the way the sample was obtained introduce (selection)bias? yes Was the sample of subjects representative with regard to the population to which the findings will be referred? unclear Was the sample size based on pre-study

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Source of funding Not reported	rancipants	Interventions	Methous	Outcomes and results	considerations of statistical power? no Was a satisfactory response rate achieved? yes Are the measurements (questionnaires) likely to be valid and reliable? yes Was the statistical significance assessed? not relevant Are confidence intervals given for the main results? no Could there be confounding factors that haven't been accounted for? yes Can the results be
					applied to your organization? yes
Full citation Ferreira, F. T., Kobayashi, E., Lopes- Cendes, I., Cendes, F., Structural abnormalities are similar in familial and nonfamilial mesial temporal lobe epilepsy, Canadian Journal of	Sample size N=67 Characteristics Age of follow up, years, mean (range): 35 (8 to 76) Syndrome type, n (%): temporal lobe epilepsy, n=67 (100)	Interventions MRI scan 2.0-t	Details Patients were recruited from the author's epilepsy clinic and all underwent the same MRI protocol	Results Proportion identified with a clinically relevant abnormality: Scarring: 2/67 Congenital/developmental: 6/67	Limitations The quality of this study was assessed using the CEBMA checklist Did the study address a clearly focused question / issue? yes Is the research method (study design)

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Neurological Sciences, 31, 368- 372, 2004	Inclusion criteria				appropriate for answering the research question? yes
Ref Id 1158443	Not reported				Is the method of selection of the subjects
Country/ies where the study was carried out	Exclusion criteria Lateral temporal lobe epilepsy				(employees, teams, divisions, organizations) clearly described? no
Brazil Study type					Could the way the sample was obtained introduce (selection)
Retrospective cohort					bias? yes
Aim of the study To assess temporal lobe structures in patients with familial temporal lobe epilepsy					Was the sample of subjects representative with regard to the population to which the findings will be referred? unclear
Study dates Not reported					Was the sample size based on pre-study considerations of
Source of funding Two of the authors					statistical power? no
received scholarship grants from Fundação de Amparo à Pesquisa do Estado					Was a satisfactory response rate achieved? yes
de São Paulo (FAPESP)					Are the measurements (questionnaires) likely to be valid and reliable? yes
					Was the statistical significance assessed? not relevant

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
					Are confidence intervals given for the main results? no Could there be confounding factors that haven't been accounted for? yes Can the results be
					applied to your organization? yes
					Other information All patients had temporal lobe epilepsy (familial and non familial)
Full citation Gaillard, W. D., Weinstein, S., Conry, J., Pearl, P. L., Fazilat, S., Vezina, L. G., Reeves-Tyer, P., Theodore, W. H., Prognosis of children with partial epilepsy: MRI and serial	Sample size N= 38 Characteristics Age at seizure onset, years, mean (range): 5.8 (0.9 to 11.9) Seizure type, n (%): partial epilepsy, 8 (100)	Interventions MRI scan 1.5-t	Details Children were referred to the epilepsy clinical and scanned using a standardised protocol. MRI imaging was interpreted by a	Results Proportion identified with a clinically relevant abnormality: Scarring: 12/38	Limitations The quality of this study was assessed using the CEBMA checklist Did the study address a clearly focused question / issue? yes Is the research method
18FDG-PET, Neurology, 68, 655- 659, 2007	Inclusion criteria Those with more than 3 partial		neuroradiologist blinded to the child's identity.		(study design) appropriate for answering the research question? yes
Ref Id 1158995	seizures before their first FDG- PET Exclusion criteria				Is the method of selection of the subjects (employees, teams,

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Country/ies where the study was carried out US Study type Retrospective cohort Aim of the study To assess the prevance of brain abnormalities in children with partial epilepsy Study dates Not reported Source of funding Not reproted	Children with a history of head trauma, meningitis, or encephalitis, and focal neurologic examinations, or benign partial epilepsy syndromes (for example, rolandic epilepsy) Those with a mass or other structural lesion (such a tumour)				divisions, organizations) clearly described? no Could the way the sample was obtained introduce (selection)bias? yes Was the sample of subjects representative with regard to the population to which the findings will be referred? unclear Was the sample size based on pre-study considerations of statistical power? no Was a satisfactory response rate achieved? yes Are the measurements (questionnaires) likely to be valid and reliable? yes Was the statistical significance assessed? not relevant Are confidence intervals given for the main results? no Could there be confounding factors that

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
					haven't been accounted for? yes Can the results be applied to your organization? Yes
Full citation Griffiths, P. D., Coley, S. C., Connolly, D. J. A., Hodgson, T., Romanowski, C. A. J., Widjaja, E., Darwent, G., Wilkinson, I. D., MR imaging of patients with localisation-related seizures: Initial experience at 3.0T and relevance to the NICE guidelines, Clinical Radiology, 60, 1090-1099, 2005 Ref Id 1086050 Country/ies where the study was carried out UK Study type Retrospective cohort Aim of the study To describe the initial experience of imaging	Sample size N=120 people with localisation related epilespsy Characteristics Age at seizure onset, years, median (range): 13 (range 25-38 years) Males, n (%): 48 (40) Seizure type, n (%): localisation related epilepsy, 120 (100) Inclusion criteria Those above 16 years-old with localisation-related epilepsy Exclusion criteria Not reported	Interventions MRI scan 3.0-t	Patients were referred to the MRI facility from a regional neuroscience centre with a new diagnosis of localisation-related epilepsy. Diagnosis was based clinically and/or electrophysiologic ally and scans were reviewed by experienced neuroradiologists.	Results Proportion identified with a clinically relevant abnormality: Tumours: 4/120 Vascular: 7/120 Scarring: 10/120 Congenital/developmental: 4/120 Inflammatory/infective/immune: 3/120	Limitations The quality of this study was assessed using the CEBMA checklist Did the study address a clearly focused question / issue? yes Is the research method (study design) appropriate for answering the research question? yes Is the method of selection of the subjects (employees, teams, divisions, organizations) clearly described? yes Could the way the sample was obtained introduce (selection)bias? yes Was the sample of subjects representative with regard to the population to which the

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
in adults with localisation-related epilepsy					findings will be referred? unclear
Study dates Not reported					Was the sample size based on pre-study considerations of statistical power? no
Source of funding Not reported					Was a satisfactory response rate achieved? yes
					Are the measurements (questionnaires) likely to be valid and reliable? yes
					Was the statistical significance assessed? not relevant
					Are confidence intervals given for the main results? no
					Could there be confounding factors that haven't been accounted for? yes
					Can the results be applied to your organization? yes
Full citation Hakami, T., McIntosh, A., Todaro, M., Lui, E., Yerra, R., Tan, K. M., French, C., Li, S.,	Sample size N= 993 adults with new-onset seizures; MRI was available in n=764	Interventions Before October 2007, MRI scans were performed	Details The first presentation to the clinic was within a median of	Results Proportion identified with a clinically relevant abnormality: 177/764 Tumours: 26/764	Limitations The quality of this study was assessed using the CEBMA checklist

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Desmond, P., Matkovic, Z., O'Brien, T. J., MRI-identified	Characteristics Age of follow up, years, mean (SD): 42.2 (18.8), range 14.3–94.3	on 1.5-tesla. From October 2007, scans	24 days (IQR 14 to 44 days) from the suspected	Vascular: 26/764 Scarring: 99/764 Congenital/developmental:	Did the study address a clearly focused question / issue? yes
pathology in adults		were performed	seizure.	26/764	,
with new-onset	Males, n (%): 597 (61)	on 3-tesla			Is the research method
seizures, Neurology,			Patients	Proportion identified with a	(study design)
81, 920-927, 2013	Seizure type, n (%): 713 (72)		presented to the	non-epilepsy related	appropriate for
	epileptic seizure [focal convulsive		clinic referred by	abnormality: 165/764	answering the research
Ref Id	in 184 patients (26), focal		their general		question? yes
1155699	nonconvulsive in 85 (12), primarily		practitioner after		
	generalized convulsive in 69 (10),		their first		Is the method of
Country/ies where	and generalized nonconvulsive in		suspected		selection of the subjects
the study was	10 (1)], 180 (18) nonepileptic event		seizure.		(employees, teams,
carried out Australia	[included syncope in 114 patients (63) and psychogenic in 66 (37)],		EEG and MRI were routinely		divisions, organizations) clearly described? yes
Australia	and 100 (10) uncertain. Seizures		requested, unless		clearly described? yes
Study type	were unclassified in 365 patients		MRI was		Could the way the
Prospective cohort	(51)		contraindicated.		sample was obtained
	(3.)				introduce
Aim of the study	Syndrome type, n (%): focal in 343		If several MRI		(selection)bias? yes
To assess the	(48), idiopathic generalized		scans were		, ,
frequency of	epilepsy (IGE) in 77 (11), and		available, the		Was the sample of
epileptogenic lesions	unclassified in 293 patients (41)		closest to the time		subjects representative
on MRI in adults with			of the last seizure		with regard to the
new-onset seizures	Previous CT: some patients did		was chosen.		population to which the
.	have previous CT at the request of				findings will be referred?
Study dates	their referring doctor. % of patients		Initially, 1		yes
January 2000 to December 2009	was not reported		neuroradiologist assessed the		Mag the comple size
December 2009			scans and a		Was the sample size based on pre-study
Source of funding	Inclusion criteria		second one		considerations of
The Royal Melbourne			assessed a		statistical power? no
Hospital	Not reported		random sample of		otation power: 110
Neuroscience	Exclusion criteria		scans.		Was a satisfactory
Foundation and by					response rate achieved?
the NHMRC Centre	 Prior diagnosis of epilepsy 		Disagreements		yes
for Research	 Those with acute symptomatic 		were resolved by		
Excellence in	seizures		a third		Are the measurements
			neuroradiologist.		(questionnaires) likely to

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Translational Neuroscience					be valid and reliable?
Neuroscience					yes
					Was the statistical significance assessed? not relevant
					Are confidence intervals given for the main results? no
					Could there be confounding factors that haven't been accounted for? no
					Can the results be applied to your organization? yes
Full citation Harini, C., Sharda, S., Bergin, A. M., Poduri, A., Yuskaitis, C. J., Peters, J. M., Rakesh,	Sample size N=71 children with infantile spasms Characteristics Age at seizure onset, years,	Interventions MRI scan 1.5 or 3-t	Details Patients were identified by searching key terms on	Proportion identified with a clinically relevant abnormality: Vascular: 15/71	Limitations The quality of this study was assessed using the CEBMA checklist
K., Kapur, K., Pearl, P. L., Prabhu, S. P., Detailed Magnetic	<u>median:</u> 6 <u>Males, n (%):</u> 31 (43.66)		institutional billing databases, inpatient and	Scarring: 4/71 Congenital/developmental: 29/71	Did the study address a clearly focused question / issue? yes
Resonance Imaging (MRI) Analysis in Infantile Spasms, Journal of Child Neurology, 33, 405- 412, 2018	Syndrome type, n (%): infantile spasms, 71 (100)		outpatient databases. Scans were interpreted by a neuroradiologist	Inflammatory/infective/imm une: 3/71	Is the research method (study design) appropriate for answering the research question? yes
	Inclusion criteria				
Ref Id 1157355	 Infants between 2 months and 2 years of age with new diagnosis of infantile spasms 				Is the method of selection of the subjects (employees, teams, divisions, organizations) clearly described? yes

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Country/ies where the study was carried out US Study type Retrospective cohort Aim of the study To describe MRI findings in children with infantile spasms Study dates January 2012 to December 2014 Source of funding No financial support for the research, authorship, and/or publication	Electroencephalographic features compatible with this diagnosis (hypsarrhythmia, modified hypsarrhythmia, or other) Exclusion criteria Those without MRI data or a single visit to the hospital where the study was conducted for a second opinion (hence lacking follow-up data) Those with infantile spasms and tuberous sclerosis complex				Could the way the sample was obtained introduce (selection)bias? yes Was the sample of subjects representative with regard to the population to which the findings will be referred? unclear Was the sample size based on pre-study considerations of statistical power? no Was a satisfactory response rate achieved? yes Are the measurements (questionnaires) likely to be valid and reliable? yes Was the statistical significance assessed? not relevant Are confidence intervals given for the main results? no Could there be confounding factors that haven't been accounted for? yes

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
					Can the results be applied to your organization? yes
Full citation Hesdorffer, D. C., Chan, S., Tian, H., Allen Hauser, W., Dayan, P., Leary, L. D., Hinton, V. J., Are MRI-detected brain abnormalities associated with febrile seizure type?, Epilepsia, 49, 765- 771, 2008 Ref Id 1159207 Country/ies where the study was carried out US Study type Prospective cohort Aim of the study To determine the yield of MRI-detected brain abnormalities in children with first febrile seizures Study dates March 1999 to April 2004	Sample size N=159 Characteristics Age at seizure onset, months (%): <18 months, n=75 (47.2); ≥18 months, n=84 (52.8) Males, n (%): 87 (54.7) Inclusion criteria • Those with first febrile seizures aged between 6 months and 5 years Exclusion criteria • Not reported	Interventions MRI scan 1.5-t	Details Children were selected by reviewing cases from the emergency department or hospital records with the ICD-9 code of 780.3 Children were classified as having febrile seizures by an epileptologist blind to the child's MRI findings and prior clinical history. MRI readings were done by a single neuroradiologist with epilwpsy expertise.	Proportion identified with a clinically relevant abnormality: Scarring: 9/159 Congenital/developmental: 9/159 Proportion identified with a non-epilepsy related abnormality: 2/159	Limitations The quality of this study was assessed using the CEBMA checklist Did the study address a clearly focused question / issue? yes Is the research method (study design) appropriate for answering the research question? yes Is the method of selection of the subjects (employees, teams, divisions, organizations) clearly described? yes Could the way the sample was obtained introduce (selection)bias? yes Was the sample of subjects representative with regard to the population to which the findings will be referred? unclear

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Source of funding National Institute of Child Health and Human Development					Was the sample size based on pre-study considerations of statistical power? no Was a satisfactory response rate achieved? yes Are the measurements (questionnaires) likely to be valid and reliable? yes Was the statistical significance assessed? not relevant Are confidence intervals given for the main results? no Could there be confounding factors that haven't been accounted for? yes Can the results be applied to your organization? yes
Full citation Hnojcikova, M., Nickels, K. C., Wetjen, N. M., Buchhalter, J. R., Raffel, C., Wirrell, E. C., EEG and neuroimaging studies	Sample size N=28 Characteristics Age at seizure onset, months, mean (SD): 9.6 (12.7)	Interventions MRI scan (magnet strenght was not reported)	Details The charts of all children who had epilepsy surgery before 60 months of age at the study's clinic were reviewed. The	Results Proportion identified with a clinically relevant abnormality: Tumours: 1/28 Scarring: 9/28 Congenital/developmental: 16/28	Limitations The quality of this study was assessed using the CEBMA checklist Did the study address a clearly focused question / issue? yes

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
in young children having epilepsy surgery, Pediatric Neurology, 43, 335-340, 2010 Ref Id 1159643 Country/ies where the study was carried out US Study type Retrospective cohort Aim of the study To evaluate the yield of MRI in children having resective epilepsy surgery before the age of 5 Study dates January 2002 to June 2009 Source of funding Not reported	Age of follow up, months, mean (SD): 28.8 (17.7) Males, n (%): 15 (54) Seizure type, n (%): partial only, n=15 (50); partial and secondarily generalised, n=2 (7); spasms only, n=4 (14); spasms + secondarily generalised, n=8 (29) Learning disability, n (%): normal, n=8 (29); mild-moderate delay, n=10 (36); severe delay, n=10 (36) Inclusion criteria • Medical intractable epilepsy before 5 years old Exclusion criteria • Children who presented with acute symptomatic seizures • Those who had corpus callosotomy without resection (those who had lesionectomy, lobectomy or multilobar resection were included)		MRI findings reported were conducted preoperatively		Is the research method (study design) appropriate for answering the research question? yes Is the method of selection of the subjects (employees, teams, divisions, organizations) clearly described? yes Could the way the sample was obtained introduce (selection)bias? yes Was the sample of subjects representative with regard to the population to which the findings will be referred? unclear Was the sample size based on pre-study considerations of statistical power? no Was a satisfactory response rate achieved? yes Are the measurements (questionnaires) likely to be valid and reliable? yes

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
					Was the statistical significance assessed? not relevant Are confidence intervals given for the main results? no Could there be confounding factors that haven't been accounted for? yes Can the results be applied to your organization? yes
Full citation Hsieh, D. T., Chang, T., Tsuchida, T. N., Vezina, L. G., Vanderver, A., Siedel, J., Brown, K., Berl, M. M., Stephens, S., Zeitchick, A., Gaillard, W. D., New-onset afebrile seizures in infants: Role of neuroimaging, Neurology, 74, 150- 156, 2010 Ref Id 1154172 Country/ies where the study was carried out US	Sample size N=317 in total, of which n=182 infants had MRI Characteristics Age of follow up: all <24 months Males, n (%): 165 (52) Seizure type, n (%): partial n=154 (48.5); no clear partial features n=163 (151.5) Learning disability, n (%): 15 (4.7) Previous CT, n (%): 298 (94) Inclusion criteria Those between 1 and 24 months Those presenting in the emergency department or as	Interventions MRI scan 1.5-t	Details MRI scans were interpreted by a paediatric neurologist. MRI sequence was the same for all the infants included. MRI was performed when focal findings were present, when CT was ambiguous or to define abnormal findings on CT	Proportion identified with a clinically relevant abnormality: Tumours: 2/182 Vascular: 24/182 Scarring: 9/182 Congenital/developmental: 51/182 Inflammatory/infective/immune: 1/182 Metabolic/genetic: 3/182 Proportion identified with a non-epilepsy related abnormality: 33/182	Limitations The quality of this study was assessed using the CEBMA checklist Did the study address a clearly focused question / issue? yes Is the research method (study design) appropriate for answering the research question? yes Is the method of selection of the subjects (employees, teams, divisions, organizations) clearly described? yes

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Study type Prospective cohort Aim of the study To assess the yield of neuroimaging in infants with new-onset afebrile seizures Study dates January 2001 to February 2007 Source of funding No specific funding was reported	inpatients in the hospital where the study was conducted with new onset afebrile seizures Exclusion criteria Those with a febrile illness Those with an infection of the CNS Those admitted for a suspicion of seizures, but discharged with a diagnosis of a "spell"	Interventions	WIGHTIOUS		Could the way the sample was obtained introduce (selection)bias? yes Was the sample of subjects representative with regard to the population to which the findings will be referred? unclear Was the sample size based on pre-study considerations of statistical power? no Was a satisfactory response rate achieved? yes Are the measurements (questionnaires) likely to be valid and reliable? yes Was the statistical significance assessed? not relevant Are confidence intervals given for the main results? no Could there be confounding factors that haven't been accounted for? yes

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
					Can the results be applied to your organization? yes
Full citation Jasim, H. A., Abdulsattar, O. A., MRI findings in iraqi patients with epilepsy: A cross sectional study, Indian Journal of Public Health Research and Development, 9, 810- 814, 2018 Ref Id 1157380 Country/ies where the study was carried out Iraq Study type Cross-sectional Aim of the study To evaluate MRI findings in patients with epilepsy Study dates 1 January 2017 to 4 June 2018 Source of funding No funding was received	Characteristics Age, years, mean (SD): 21.31 (12.75) Males, n (%): 26 (50.9) Seizure type, n (%): focal: 36 (70.6); generalised: 15 (29.4) Inclusion criteria Not reported Exclusion criteria Children with a history of acute cerebral insult, such as infection, trauma, metabolic abnormalities or vascular pathology. Those with neonatal seizures were also excluded	Interventions MRI 1.5 t	Details Patients were referred to the neurology department of the hospital where the study took place. MRI protocol was the same for all patients.	Results Clinically relevant abnormalities: Tumours: 6/51 Scarring: 11/51	Limitations The quality of this study was assessed using the CEBMA checklist Did the study address a clearly focused question / issue? yes Is the research method (study design) appropriate for answering the research question? yes Is the method of selection of the subjects (employees, teams, divisions, organizations) clearly described? no Could the way the sample was obtained introduce (selection)bias? yes Was the sample of subjects representative with regard to the population to which the findings will be referred? unclear Was the sample size based on pre-study

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Study details	ranticipants	interventions	Wetflous	Outcomes and Results	considerations of statistical power? no Was a satisfactory response rate achieved? yes Are the measurements (questionnaires) likely to be valid and reliable? yes Was the statistical significance assessed? not relevant Are confidence intervals given for the main results? no Could there be confounding factors that haven't been accounted for? yes Can the results be applied to your organization? yes
Full citation Jeniffer, V. N., Udayakumar, S., Pushpalatha, K., A clinical study to identify the possible etiology of complex partial seizures using magnetic resonance imaging brain findings	Sample size N=64 Characteristics Age of follow up, years: all <18 years old; results have not been reported separately by age Males, n (%): 42 (65.6)	Interventions MRI scan 1.5-t	Details A detailed clinical evaluation was carried out in all children, which included blood tests and MRI scan. MRI protocol was the	Results Proportion identified with a clinically relevant abnormality: Tumours: 1/64 Scarring: 10/64 Congenital/developmental: 29/64	Limitations The quality of this study was assessed using the CEBMA checklist Did the study address a clearly focused question / issue? yes

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
and its implications on treatment, Journal of Pediatric Neurosciences, 10, 350-354, 2015 Ref Id 1156379 Country/ies where the study was carried out India Study type Prospective cohort Aim of the study To assess MRI findings in children aged 1 to 12 years old with complex partial seizures Study dates October 2011 to	Participants Learning disability, n (%): 0 (0) Inclusion criteria Those aged between 1 and 18 years old Those diagnosed with complex partial seizures Those attending the department of paediatrics where the sutudy was conducted Those who gave consent to participate Exclusion criteria Those with developmental delay, learning disabilities or cerebral palsy Those with seizures following head injury	Interventions	Methods same for all children.	Outcomes and Results	Is the research method (study design) appropriate for answering the research question? yes Is the method of selection of the subjects (employees, teams, divisions, organizations) clearly described? no Could the way the sample was obtained introduce (selection)bias? yes Was the sample of subjects representative with regard to the population to which the findings will be referred? unclear Was the sample size based on pre-study
					based on pre-study considerations of
Source of funding No funding was received					statistical power? no Was a satisfactory response rate achieved? yes
					Are the measurements (questionnaires) likely to be valid and reliable? yes

Full citation Koirala, K., Magnetic resonance neuroimaging in patient with complain of seizure, Journal of Nepal Health Research Council, 9, 56-60, 2011 Ref Id 1159895 Country/les where the study was carried out Nepal Country/les where the study was carried out Nepal Exclusion criteria Linterventions MRI scan 0.2-t Interventions MRI scan 0.2-t Interventions MRI scan 0.2-t All patients underwent the same MRI protocol. No further details were provided was assessed using the CEBMA checklist were provided scaring; 6/160 Congenital/developmental: 1/160 Inflammatory/infective/imm une: 12/160 Is the research method (study design) appropriate for answering the research question of selection of the subjects (employees, teams,	Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Full citation Koirala, K., Magnetic resonance neuroimaging in patient with complain of seizure, Journal of Nepal Health Research Council, 9, 56-60, 2011 Ref Id 1159895 Country/les where the study was carried out Nepal Study type Sample size N=160 Interventions MRI scan 0.2-t MRI scan						significance assessed? not relevant Are confidence intervals given for the main results? no
Koirala, K., Magnetic resonance neuroimaging in patient with complain of seizure, Journal of Nepal Health Research Council, 9, 56-60, 2011 Ref Id 1159895 Country/ies where the study was carried out Nepal N=160 MRI scan 0.2-t MRI scan 0.2-t All patients underwent the same MRI protocol. No further details were provided MRI scan 0.2-t All patients underwent the same MRI protocol. No further details were provided MRI scan 0.2-t All patients underwent the same MRI protocol. No further details were provided Scarring: 6/160 Congenital/developmental: 1/160 Inclusion criteria • The quality of this study was assessed using the CEBMA checklist Tumours: 21/160 Did the study address a clearly focused question / issue? yes Inclusion criteria • Those diagnosed with epilepsy and referred to a private epilepsy clinic to perform a MRI within 1 year Study type Study type						Can the results be applied to your
Aim of the study	Koirala, K., Magnetic resonance neuroimaging in patient with complain of seizure, Journal of Nepal Health Research Council, 9, 56-60, 2011 Ref Id 1159895 Country/ies where the study was carried out Nepal Study type Cross-sectional	N=160 Characteristics Age of follow up, years, n (%): 1 to 82 years old; n=36 (22.5) were ≥16 years old; n=124 (77.5) were >16 years old Inclusion criteria • Those diagnosed with epilepsy and referred to a private epilepsy clinic to perform a MRI within 1 year Exclusion criteria		All patients underwent the same MRI protocol. No further details	Proportion identified with a clinically relevant abnormality: Tumours: 21/160 Vascular: 11/160 Scarring: 6/160 Congenital/developmental: 1/160 Inflammatory/infective/imm	The quality of this study was assessed using the CEBMA checklist Did the study address a clearly focused question / issue? yes Is the research method (study design) appropriate for answering the research question? yes Is the method of selection of the subjects (employees, teams, divisions, organizations)

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
To assess the yield of MRI abnormalities in patients with epilepsy Study dates					Could the way the sample was obtained introduce (selection)bias? yes
July 2008 to June 2009 Source of funding Not reported					Was the sample of subjects representative with regard to the population to which the findings will be referred? unclear
					Was the sample size based on pre-study considerations of statistical power? no
					Was a satisfactory response rate achieved? yes
					Are the measurements (questionnaires) likely to be valid and reliable? yes
					Was the statistical significance assessed? not relevant
					Are confidence intervals given for the main results? no
					Could there be confounding factors that haven't been accounted for? yes

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
					Can the results be applied to your organization? yes
Full citation Labate, A., Ventura, P., Gambardella, A., Le Piane, E., Colosimo, E., Leggio, U., Ambrosio, R., Condino, F., Messina, D., Lanza, P., Aguglia, U., Quattrone, A., MRI evidence of mesial temporal sclerosis in sporadic "benign" temporal lobe epilepsy, Neurology, 66, 562-565, 2006 Ref Id 1158857	Sample size N=101 people with sporadic benign temporal lobe epilepsy Characteristics Age at seizure onset, years, mean (SD): 22.3 (17.4) Age of follow up, years, mean (SD): 37.3 (17.5) Males, n (%): 50 (49.5) Seizure type: people were either seizure free, had auras, or not more than 2 disabling seizures per year for at least 2 years (with or without appropriate antiepileptic medication)	Interventions MRI scans performed on a 1.5-tesla	Details In each person, the diagnosis of temporal lobe epilepsy was made on the basis of clinical, EEG and MRI criteria. All patients had MRI evaluations based on a protocol routinely used for those with temporal lobe epilepsy.	Results Proportion identified with a clinically relevant abnormality: 39/101 Scarring: 39/101	Limitations The quality of this study was assessed using the CEBMA checklist Did the study address a clearly focused question / issue? yes Is the research method (study design) appropriate for answering the research question? yes Is the method of selection of the subjects (employees, teams, divisions, organizations) clearly described? yes
Country/ies where the study was carried out Italy	Syndrome type: sporadic benign temporal lobe epilepsy				Could the way the sample was obtained introduce (selection)bias? yes
Study type Retrospective cohort Aim of the study To assess whether there is MRI- detectable mesial	Inclusion criteria Not reported Exclusion criteria Any suggestion of seizure onset				Was the sample of subjects representative with regard to the population to which the findings will be referred? unclear
temporal sclerosis in people with sporadic	outside the mesial temporal structures by semiology or EEG findings				Was the sample size based on pre-study

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
benign temporal lobe epilepsy					considerations of statistical power? no
Study dates Not reported					Was a satisfactory response rate achieved? yes
Source of funding Not reported					Are the measurements (questionnaires) likely to be valid and reliable? yes
					Was the statistical significance assessed? not relevant
					Are confidence intervals given for the main results? no
					Could there be confounding factors that haven't been accounted for? no
					Can the results be applied to your organization? yes
Full citation Lefkopoulos, A., Haritanti, A., Papadopoulou, E., Karanikolas, D., Fotiadis, N., Dimitriadis, A. S., Magnetic resonance imaging in 120 patients with	Sample size N=120 people with intractable partial seizures Characteristics Age of follow up, years, mean (SD): 21 (SD not reported) Males, n (%): 40 (33.3)	Interventions MRI scan 1.5-t	Details Not reported	Results Proportion identified with a clinically relevant abnormality: Vascular: 9/120 Scarring: 30/120 Congenital/developmental: 23/120 Inflammatory/infective/immune: 4/120	Limitations The quality of this study was assessed using the CEBMA checklist Did the study address a clearly focused question / issue? yes

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
intractable partial seizures: A preoperative assessment, Neuroradiology, 47, 352-361, 2005	Seizure type, n(%): intractable partial, 120 (100) Response to treatment: existing diagnosis and treatment resistant, 120 (100)				Is the research method (study design) appropriate for answering the research question? yes
Ref Id 1158669					Is the method of selection of the subjects (employees, teams, divisions, organizations)
Country/ies where	Inclusion criteria				clearly described? no
the study was carried out Greece	 Those with intractable partial seizures 				Could the way the sample was obtained
Study type Retrospective cohort	Not reported				introduce (selection)bias? unclear (how the sample was obtained was not
Aim of the study					reported)
To assess MRI findings in people with intractable partial seizures					Was the sample of subjects representative with regard to the population to which the
Study dates January 2000 to June 2003					findings will be referred? unclear (as above)
					Was the sample size based on pre-study
Source of funding Not reported					considerations of statistical power? no information was provided
					Was a satisfactory response rate achieved? yes

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
					Are the measurements (questionnaires) likely to be valid and reliable? yes Was the statistical significance assessed? not applicable Are confidence intervals given for the main results? no Could there be confounding factors that haven't been accounted for? no Can the results be applied to your organization? yes
Full citation Ma, W., Li, C., Liu, L., Li, S., Liu, Y., Pre- Operative Interictal Discharge Patterns and Magnetic Resonance Imaging Findings Affect Prognosis of Temporal Lobe Epilepsy Surgery, European Neurology, 81, 152-162, 2019 Ref Id 1157748	Sample size N=115 Characteristics Age of follow up, years, mean (SD): 30.8 (12.6) Males, n (%): 59 (51.3) Seizure type, n (%): 115 (100) temporal lobe epilepsy Response to treatment, n (%): 115 (100) existing diagnosis and treatment resistant Inclusion criteria	Interventions MRI scan (strength of magnet was not reported)	Details Participants were attending the neurosurgery department of the hospital where the study was conducted. Diagnosis was made on the basis of clinical presentation and EEG monitoring	Results Proportion identified with a clinically relevant abnormality: Tumours: 18/115 Vascular: 7/115 Scarring: 42/115 Congenital/developmental: 5/115 Inflammatory/infective/immune: 8/115	Limitations The quality of this study was assessed using the CEBMA checklist Did the study address a clearly focused question / issue? yes Is the research method (study design) appropriate for answering the research question? yes Is the method of selection of the subjects

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Country/ies where the study was carried out China Study type Retrospective cohort Aim of the study To assess MRI findings in people with temporal lobe epilepsy Study dates October 2010 to October 2014 Source of funding No specific grant or funding was received to conduct this study	Patients attending the neurosurgery department of the hospital where the study was conducted and presenting with temporal lobe epilepsy Exclusion criteria Not reported	IIILEI VEITUOIIS	Wethous	Outcomes and Results	(employees, teams, divisions, organizations) clearly described? no Could the way the sample was obtained introduce (selection) bias? yes Was the sample of subjects representative with regard to the population to which the findings will be referred? unclear Was the sample size based on pre-study considerations of statistical power? no Was a satisfactory response rate achieved? yes Are the measurements (questionnaires) likely to be valid and reliable? yes Was the statistical significance assessed? not applicable Are confidence intervals given for the main results? no

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
					Could there be confounding factors that haven't been accounted for? yes Can the results be applied to your organization? yes
Full citation Nair, P. P., Kalita, J., Misra, U. K., Role of cranial imaging in epileptic status, European Journal of Radiology, 70, 475- 80, 2009 Ref Id 1154726 Country/ies where the study was carried out India Study type Prospective cohort Aim of the study To assess the role of imaging in predicting the outcome of status epilepticus Study dates January 2002 to March 2007	Sample size N=99 people with status epilepticus of which n=41 underwent MRI Characteristics Age of follow up, years, mean (range): 35 (1 to 78) Males, n (%): 59 (59) Seizure type, n (%): 99 (100) status epilepticus Previous CT, n (%): MRI and CT was carried out in n=14 (14) Inclusion criteria Those diagnosed with status epilepticus and attending the emergency department of the hospital where the study was carried out Those developing status epilepticus during their hospital stay in the neurology department of the hospital where the study was carried out Exclusion criteria Those with pseudoseizures	Interventions MRI scan 1.5-t	Details A detailed clinical examination was conducted for all patients. Status epilepticus was defined as the occurrence of 2 or more seizures without full recovery of consciousness between the seizures, or continuous convulsive activity for >10 minutes.	Results Proportion identified with a clinically relevant abnormality: Vascular: 4/41 Inflammatory/infective/imm une: 35/41	Limitations The quality of this study was assessed using the CEBMA checklist Did the study address a clearly focused question / issue? yes Is the research method (study design) appropriate for answering the research question? yes Is the method of selection of the subjects (employees, teams, divisions, organizations) clearly described? no Could the way the sample was obtained introduce (selection)bias? yes Was the sample of subjects representative with regard to the population to which the

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Source of funding Not reported					findings will be referred? unclear
					Was the sample size based on pre-study considerations of statistical power? no
					Was a satisfactory response rate achieved? yes
					Are the measurements (questionnaires) likely to be valid and reliable? yes
					Was the statistical significance assessed? not relevant
					Are confidence intervals given for the main results? no
					Could there be confounding factors that haven't been accounted for? yes
					Can the results be applied to your organization? yes
Full citation Petrou, M., Foerster, B., Maly, P. V.,	Sample size N=437	Interventions MRI scan (strength of	Details MRI imaging was performed as part	Results Proportion identified with a clinically relevant	Limitations The quality of this study was assessed using
Eldevik, O. P., Leber, S., Sundgren, P. C.,	Characteristics	magnet was not reported)	of an initial seizure workup.	<u>abnormality:</u> Tumours: 4/437	the CEBMA checklist

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
					be valid and reliable? yes Was the statistical significance assessed? not applicable Are confidence intervals given for the main results? no Could there be confounding factors that haven't been accounted for? yes Can the results be applied to your organization? yes
Full citation Rasool, A., Choh, S. A., Wani, N. A., Mushtaq Ahmad, S., Iqbal, Q., Role of electroencephalogram and neuroimaging in first onset afebrile and complex febrile seizures in children from Kashmir, Journal of Pediatric Neurosciences, 7, 9- 15, 2012 Ref Id 1154932	Sample size N=276, of which n=157 received MRI Characteristics Age of follow up, range: 6 months to 14 years old Males, n (%): 162 (58.7) Seizure type, n (%): partial, n= 86 (31.1); generalised, n=116 (42); complex febrile seizures, n= 64 (23); undetermined, n=10 (3.6) Learning disability, n (%): 0 (0) Inclusion criteria • Not reported	Interventions MRI scan 1.5-t	Details Participants were patients attending the emergency, inpatients, or outpatient departments of advanced paediatrics. The International League Against Epilepsy classification was used to define seizure types.	Results Proportion identified with a clinically relevant abnormality: Scarring: 2/157 Congenital/developmental: 9/157 Proportion identified with a non-epilepsy related abnormality: 4/157	Limitations The quality of this study was assessed using the CEBMA checklist Did the study address a clearly focused question / issue? yes Is the research method (study design) appropriate for answering the research question? yes Is the method of selection of the subjects (employees, teams,

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Country/ies where the study was carried out India Study type Prospective cohort Aim of the study To assess the frequency of abnormal neuroimaging in children with newonset afebrile and febrile seizures Study dates November 2006 to November 2008 Source of funding No funding was received	Exclusion criteria Those with seizures resulting from an acute situational etiology (for example, toxin infection, trauma) Those with a chronic neurologic illness (for example, cerebral palsy, learning disabilities, pervasive developmental disorders) Those with other abnormalities on neurologic examination or with simple febrile seizures				divisions, organizations) clearly described? yes Could the way the sample was obtained introduce (selection)bias? yes Was the sample of subjects representative with regard to the population to which the findings will be referred? unclear Was the sample size based on pre-study considerations of statistical power? no Was a satisfactory response rate achieved? yes Are the measurements (questionnaires) likely to be valid and reliable? yes Was the statistical significance assessed? not relevant Are confidence intervals given for the main results? no Could there be confounding factors that

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
					haven't been accounted for? yes Can the results be applied to your organization? yes
Full citation Santos, S. L. M., Ghizoni, E., Li, L. M., Cendes, F., Dynamic assessment of high- resolution MRI with multi-planar reconstruction increases the yield of lesion detection in patients with partial epilepsy, Journal of Epilepsy and Clinical Neurophysiology, 11, 111-116, 2005 Ref Id 1158708 Country/ies where the study was carried out Brazil Study type Retrospective cohort Aim of the study To evaluate the presence and type of lesions associated with partial epilepsy	Characteristics Age at seizure onset, years, mean (SD): 8.5 (3.1) Age of follow up, years, mean (SD): 23.9 (9) Seizure type, n (%): partial epilepsy, 100 (100) Inclusion criteria • Not reported Exclusion criteria • Not reported	Interventions MRI scan (strength magnet not reported)	Details Patients were recruited consecutively. Partial epilepsy diagnosis was based on previous EEG examinations and were established according to ILAE criteria.	Results Proportion identified with a clinically relevant abnormality: Tumours: 1/100 Vascular: 1/100 Scarring: 66/100 Congenital/developmental: 16/100 Inflammatory/infective/immune: 3/100	Limitations The quality of this study was assessed using the CEBMA checklist Did the study address a clearly focused question / issue? yes Is the research method (study design) appropriate for answering the research question? yes Is the method of selection of the subjects (employees, teams, divisions, organizations) clearly described? no Could the way the sample was obtained introduce (selection) bias? yes Was the sample of subjects representative with regard to the population to which the findings will be referred? unclear

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Study dates April to September 2008					Was the sample size based on pre-study considerations of statistical power? no
Source of funding Not reported					Was a satisfactory response rate achieved? yes
					Are the measurements (questionnaires) likely to be valid and reliable? yes
					Was the statistical significance assessed? not applicable
					Are confidence intervals given for the main results? no
					Could there be confounding factors that haven't been accounted for? yes
					Can the results be applied to your organization? yes
Full citation Sinha, S., Satishchandra, P., Kalband, B. R., Bharath, R. D., Thennarasu, K.,	Sample size N=201; n=43 patients underwent MRI Characteristics Age at seizure onset, years, mean	Interventions MRI scan 1.5-t	Details All patients underwent a detailed clinical evaluation. All patients	Results Proportion identified with a clinically relevant abnormality: Tumours: 5/43 Vascular: 13/43	Limitations The quality of this study was assessed using the CEBMA checklist
Neuroimaging	(SD): 68 (7.5)		underwent CT,	Scarring: 1/43	

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
observations in a cohort of elderly manifesting with new onset seizures: Experience from a university hospital, Annals of Indian Academy of Neurology, 15, 273-280, 2012 Ref Id 1155182 Country/ies where the study was carried out India Study type Prospective study Aim of the study To assess the MRI observations in elderly people manifesting with new onset seizures Study dates January 2007 to January 2009 Source of funding No funding was received to conduct this study	Males, n (%): 131 (65.2) Seizure type, n (%): simple partial seizure, n= 84 (42); generalised tonic clonic seizure, n=61 (30.3); complex partial seizure, n=55 (27.4) Syndrome type, n (%): acute symptomatic, n=86 (42.3); remote symptomatic, n=37 (18.4); cryptogenic, n=75 (37.8); idiopathic, n=3 (1.5) Previous CT, n (%): 201 (100) Inclusion criteria Those who manifested with new onset seizures in the neurology department of the hospital where the study was conducted who manifested with new onset seizures Exclusion criteria Those with epilepsy and onset before 60 years old		and only those in whom it was clinically indicated had a MRI scan	Inflammatory/infective/imm une: 5/43	Did the study address a clearly focused question / issue? yes Is the research method (study design) appropriate for answering the research question? yes Is the method of selection of the subjects (employees, teams, divisions, organizations) clearly described? yes Could the way the sample was obtained introduce (selection)bias? yes Was the sample of subjects representative with regard to the population to which the findings will be referred? unclear Was the sample size based on pre-study considerations of statistical power? no Was a satisfactory response rate achieved? yes Are the measurements (questionnaires) likely to

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
					be valid and reliable?
					yes
					Was the statistical significance assessed? not relevant
					Are confidence intervals given for the main results? no
					Could there be confounding factors that haven't been accounted for? yes
					Can the results be applied to your organization? yes
Full citation Solosrungruang, A., Laothamatas, J., Chinwarun, Y.,	Sample size N=91 adult patients with epilepsy Characteristics	Interventions MRI scan 1.5-t	Details MRI scans were reviewer by a neuroradiologist	Proportion identified with a clinically relevant abnormality:	Limitations The quality of this study was assessed using the CEBMA checklist
Magnetic resonance imaging of the brain in epileptic adult patients: experience in Ramathibodi	Age of follow up, years, mean (range): 36.9 (15-85) Males, n (%): 37 (40.6)		or radiologist. The same MRI protocol was applied to all patients.	Tumours: 7/91 Vascular: 17/91 Scarring: 31/91 Congenital/developmental: 19/91	Did the study address a clearly focused question / issue? yes
Hospital, Journal of the Medical Association of Thailand = Chotmaihet	Syndrome type, n (%): generalised seizure, n=50 (41.67); partial seizure, n=70 (58.33) (*n=25 had their symptoms classified as more than 1 seizure type)		panomo.	Inflammatory/infective/imm une: 9/91	Is the research method (study design) appropriate for answering the research question? yes
thangphaet, 90, 762-773, 2007	Inclusion criteria				Is the method of selection of the subjects
Ref Id 1159098	Those ≥15 years old with epilepsy or seizure who had an				(employees, teams,

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Country/ies where the study was carried out	MRI scan in the hospital where the study was carried out				divisions, organizations) clearly described? yes Could the way the
Thailand Study type	 Exclusion criteria Those with incomplete MRI study and clinical data 				sample was obtained introduce (selection)bias? yes
Aim of the study To classify the imaging of structural abnormalities of epileptic adult patients referred for MRI					Was the sample of subjects representative with regard to the population to which the findings will be referred? unclear
Study dates January 2001 to December 2002					Was the sample size based on pre-study considerations of statistical power? no
Source of funding Not reported					Was a satisfactory response rate achieved? yes
					Are the measurements (questionnaires) likely to be valid and reliable? yes
					Was the statistical significance assessed? not relevant
					Are confidence intervals given for the main results? no
					Could there be confounding factors that

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
					haven't been accounted for? yes Can the results be applied to your organization? yes
Full citation Toledo, M., Sarria- Estrada, S., Quintana, M., Auger, C., Salas- Puig, X., Santamarina, E., Vert, C., Rovira, A., 3 TESLA MR imaging in adults with focal onset epilepsy, Clinical Neurology and Neurosurgery, 115, 2111-2116, 2013 Ref Id 1155884 Country/ies where the study was carried out Spain Study type Prospective cohort Aim of the study To evaluate the yield of MRI for detecting epileptogenic cerebral lesions Study dates	Sample size N=161 Characteristics Age of follow up, years, mean (SD): 41.6 (16.3) Males, n (%): 78 (64.4) Seizure type, n (%): focal, n=161 (100) Response to treatment, n (%): drug resistant, n=90 (56) Inclusion criteria Those ≥16 years old diagnosed with focal epilepsy Exclusion criteria Those with multifocal, generalized, non-classifiable, or non-epileptic seizures Those with lack of diagnostic consensus Those with multifocal or generalised epilepsy and the presence of non-epileptic seizures	Interventions MRI scan 3-t	Details Diagnosis was based on the results of clinical, MR imaging and video-EEG findings. Patients meeting inclusion criteria from the epilepsy unit of the tertiary hospital where the study was conducted where included. The diagnosis of focal epilepsy was independently established by 3 expert epileptologists	Results Proportion identified with a clinically relevant abnormality: Tumours: 17/161 Vascular: 15/161 Scarring: 27/161 Congenital/developmental: 18/161	Limitations The quality of this study was assessed using the CEBMA checklist Did the study address a clearly focused question / issue? yes Is the research method (study design) appropriate for answering the research question? yes Is the method of selection of the subjects (employees, teams, divisions, organizations) clearly described? yes Could the way the sample was obtained introduce (selection)bias? yes Was the sample of subjects representative with regard to the population to which the findings will be referred? unclear

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Not reported Source of funding Not reported					Was the sample size based on pre-study considerations of statistical power? no
					Was a satisfactory response rate achieved? yes
					Are the measurements (questionnaires) likely to be valid and reliable? yes
					Was the statistical significance assessed? not relevant
					Are confidence intervals given for the main results? no
					Could there be confounding factors that haven't been accounted for? yes
					Can the results be applied to your organization? yes
Full citation Wieshmann, U. C., Clinical application of neuroimaging in epilepsy, Journal of	Sample size N=528 people had a scan, n=495 scans were reviewed, n=332 had a MRI scan	Interventions MRI scan (standard MRI and high resolution MRI)	Details MRI scans were reviewed and imaging modality identified. The	Results Proportion identified with a clinically relevant abnormality: Tumours: 21/332	Limitations The quality of this study was assessed using the CEBMA checklist
Neurology Neurosurgery and	Characteristics	resolution wiki)	neuroradiological findings were	Vascular: 14/332 Scarring: 134/332	

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Psychiatry, 74, 466-470, 2003 Ref Id 1155495 Country/ies where the study was carried out UK Study type Cross-sectional Aim of the study To evaluate the prevalence of detected structural abnormalities in patients with epilepsy Study dates Not reported Source of funding Not reported	Age of follow up, years, mean (SD): 39.7 (14.2) Inclusion criteria Those with chronic active epilepsy, a single epileptic seizure, epilepsy in remission (no seizures for two years or longer) or nonepileptic seizures. Exclusion criteria Not reported		defined as normal, consistent with hippocampal sclerosis, vascular abnormality, tumour, malformation of cortical developlopment, brain damage, or non-specific abnormality	Congenital/developmental: 13/332	Did the study address a clearly focused question / issue? yes Is the research method (study design) appropriate for answering the research question? yes Is the method of selection of the subjects (employees, teams, divisions, organizations) clearly described? yes Could the way the sample was obtained introduce (selection) bias? yes Was the sample of subjects representative with regard to the population to which the findings will be referred? yes Was the sample size based on pre-study considerations of statistical power? no Was a satisfactory response rate achieved? yes Are the measurements (questionnaires) likely to

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
					be valid and reliable? yes
					Was the statistical significance assessed? not relevant
					Are confidence intervals given for the main results? no
					Could there be confounding factors that haven't been accounted for? no
					Can the results be applied to your organization? yes
Full citation Wongladarom, S., Laothamatas, J., Visudtibhan, A.,	Sample size N=100 children Characteristics	Interventions Scans were performed with MRI 1.5-t	Details Diagnosis was established according to	Results Proportion identified with a clinically relevant abnormality: 741/100	Limitations The quality of this study was assessed using the CEBMA checklist
Sawatsut, P., Magnetic resonance imaging of the brain in epileptic pediatric	Age of follow up, years, mean (SD): 7 (5 months) Males, n (%): 43 (43)		clinical presentation and EEG MRI was	Tumours: 3/100 Primarily generalised: 0/16 Partial: 3/26	Did the study address a clearly focused question / issue? yes
patients: Review of the experience in Ramathibodi Hospital,	Seizure type, n (%): 16 (16) children with primary generalized		performed according to a pre-specified	Complex partial seizures: 0/9 Focal with secondarily: 0/44	Is the research method (study design) appropriate for
Journal of the Medical Association of Thailand, 87, 1092-	seizure, 79 (79) children with partial or complex partial seizures with or without secondary		protocol	Infantile spasms: 0/2 Lennox-Gastaut syndrome: 0/2	answering the research question? yes
1099, 2004	generalization. The remaining 5 (5) children had a specific syndrome			Londau-Kleffner syndrome: 0/1	Is the method of selection of the subjects
Ref Id 1158559	Syndrome type, n (%): 2 (2) infantile spasms, 2 (2) Lennox-			Vascular: 5/100 Primarily generalised: 1/16 Partial: 3/26	(employees, teams, divisions, organizations) clearly described? yes

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
Country/ies where the study was carried out Thailand Study type Retrospective cohort Aim of the study To assess the MRI findings in a group of children referred with epilepsy Study dates January 1999 to December 2002 Source of funding Not reported	Gastaut Syndrome, 5 (5) Londau-Kleffner syndrome Inclusion criteria Those <15 years old Those with epilepsy or seizure and had MRI studies at the study Hospital between 1st January 1999 and 31st December 2002 Exclusion criteria Those with unavailable MRI studies and incomplete clinical data Those without evidence of seizure or epilepsy from the clinical review			Complex partial seizures: 0/9 Focal with secondarily: 1/44 Infantile spasms: 0/2 Lennox-Gastaut syndrome: 0/1 Londau-Kleffner syndrome: 0/1 Scarring: 42/100 Primarily generalised: 9/16 Partial: 6/26 Complex partial seizures: 5/9 Focal with secondarily: 19/44 Infantile spasms: 1/2 Lennox-Gastaut syndrome: 2/2 Londau-Kleffner syndrome: 0/1 Congenital/developmental: 34/100 Primarily generalised: 2/16 Partial: 8/26 Complex partial seizures: 4/26 Focal with secondarily: 18/44 Infantile spasms: 1/2 Lennox-Gastaut syndrome: 0/2 Londau-Kleffner syndrome: 1/1 Inflammatory/infective/ immune: 7/100 Primarily generalised: 2/16	Could the way the sample was obtained introduce (selection)bias? potentially, all MRI examinations were done in the same hospital Was the sample of subjects representative with regard to the population to which the findings will be referred? unclear Was the sample size based on pre-study considerations of statistical power? no Was a satisfactory response rate achieved? yes Are the measurements (questionnaires) likely to be valid and reliable? yes Was the statistical significance assessed? not applicable Are confidence intervals given for the main results? no Could there be confounding factors that

Study details	Participants	Interventions	Methods	Outcomes and Results	Comments
				Partial: 3/26 Complex partial seizures: 0/9 Focal with secondarily: 2/44 Infantile spasms: 0/2 Lennox-Gastaut syndrome: 0/2 Londau-Kleffner syndrome: 0/1 *17/100 had more than MTS in combination with other abnormality, which has been included in the scarring group Proportion identified with a non-epilepsy related abnormality: 9/100 Primarily generalised: 2/16 Partial: 3/26 Complex partial seizures: 0/9 Focal with secondarily: 4/44 Infantile spasms: 0/2 Lennox-Gastaut syndrome: 0/2 Londau-Kleffner syndrome: 0/1	haven't been accounted for? no Can the results be applied to your organization? yes

Appendix E - Forest plots

Forest plots for review question: What is the yield of relevant abnormalities detected by MRI in people with epilepsy?

This section includes forest plots only for outcomes that are meta-analysed. Outcomes from single studies are not presented here, but the quality assessment for these outcomes is provided in the GRADE profiles in appendix F.

Critical outcomes: proportion identified with tumour abnormalites

Figure 2: Proportion identified with tumour abnormalities: overall estimate

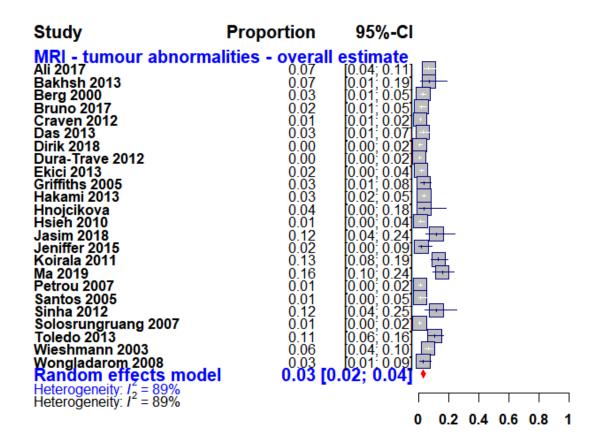


Figure 3: Proportion of tumour abnormalities identified in infants (<3 years old at seizure onset)

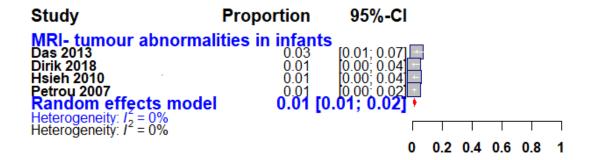


Figure 4: Proportion of tumour abnormalities identified in children (3 to 11 years old at seizure onset)

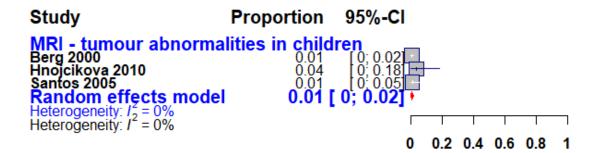


Figure 5: Proportion of tumour abnormalities identified in focal (partial) epilepsy

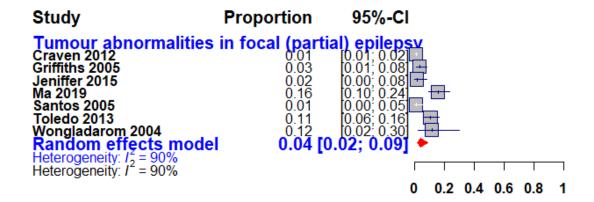


Figure 6: Proportion of tumour abnormalities identified in genetic (idiopathic) generalised epilepsy

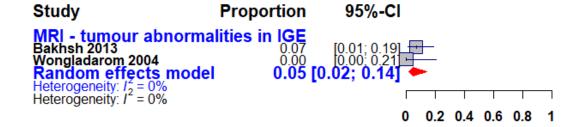


Figure 7: Proportion of tumour abnormalities identified on 1.5-t

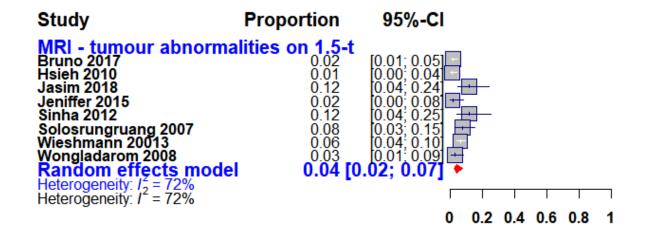


Figure 8: Proportion of tumour abnormalities identified on 3.0-t

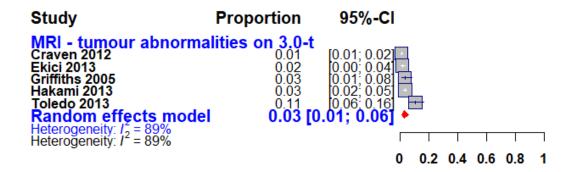


Figure 9: Proportion of tumour abnormalities identified in those with a new diagnosis

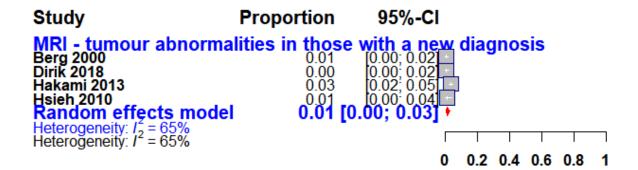


Figure 10: Proportion of tumour abnormalities identified in those with existing diagnosis and treatment resistant

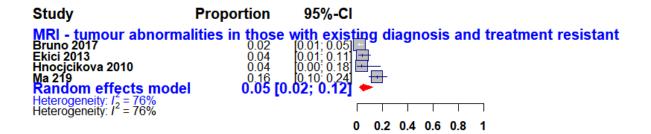
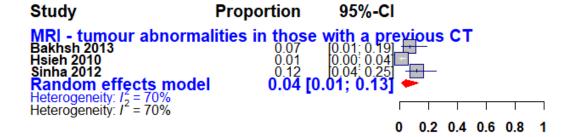


Figure 11: Proportion of tumour abnormalities identified in those with a previous CT scan



Critical outcomes: proportion identified with vascular abnormalities

Figure 12: Proportion identified with vascular abnormalities: overall estimate

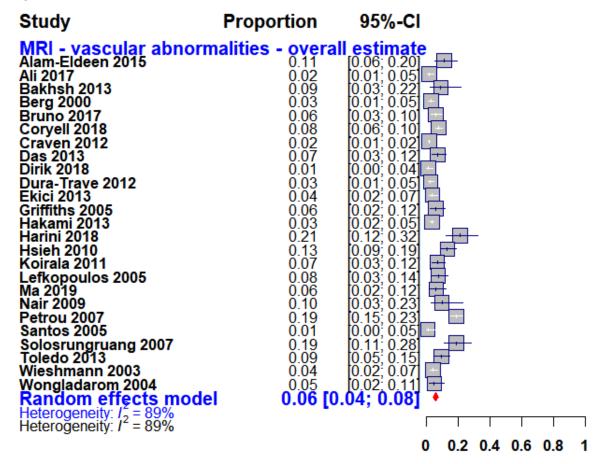


Figure 13: Proportion of vascular abnormalities identified in children (3 to 11 years old at seizure onset)

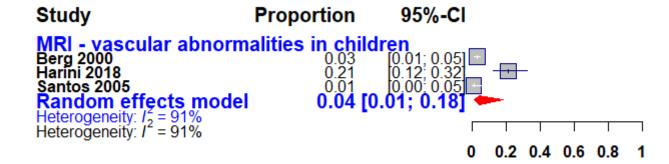


Figure 14: Proportion of vascular abnormalities identified in young people (11 to 25 years old at seizure onset)

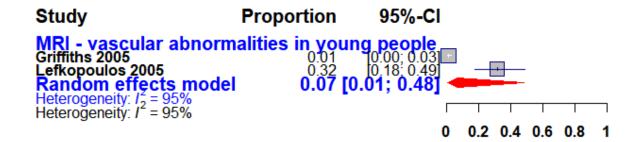


Figure 15: Proportion of vascular abnormalities identified in focal (partial) epilepsy

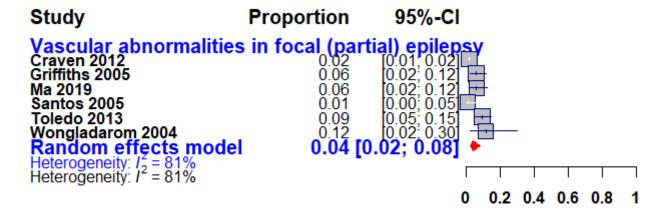


Figure 16: Proportion of vascular abnormalities identified in genetic (idiopathic) generalised epilepsy

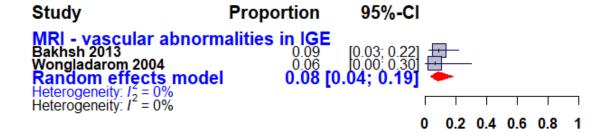


Figure 17: Proportion of vascular abnormalities identified in West syndrome

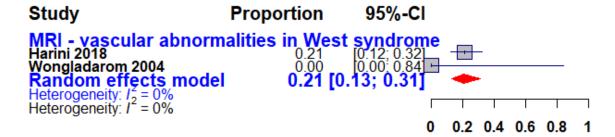


Figure 18: Proportion of vascular abnormalities identified on 1.5-t

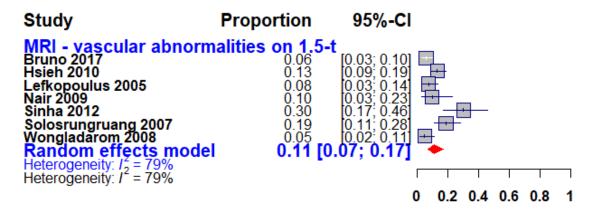


Figure 19: Proportion of vascular abnormalities identified on 3.0-t

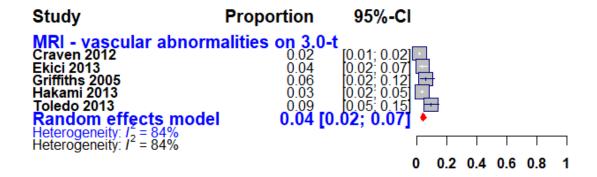


Figure 20: Proportion of vascular abnormalities identified in those with a new diagnosis

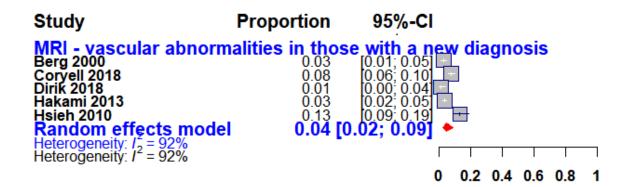
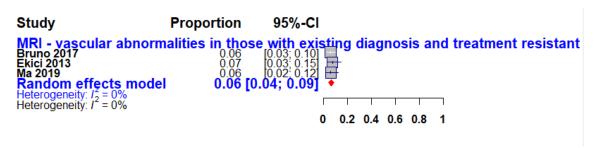


Figure 21: Proportion of vascular abnormalities identified in those with existing diagnosis and treatment resistant



Critical outcomes: proportion identified with scarring abnormalities

Figure 22: Proportion identified with scarring abnormalities: overall estimate

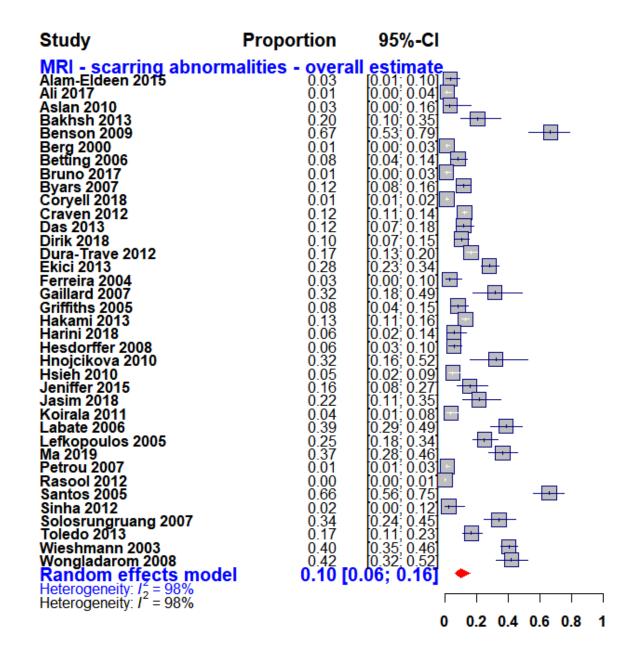


Figure 23: Proportion of scarring abnormalities identified in infants (<3 years old at seizure onset)

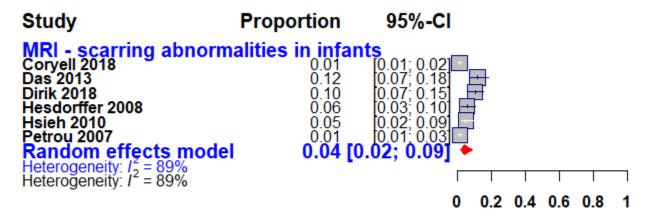


Figure 24: Proportion of scarring abnormalities identified in children (3 to 11 years old at seizure onset)

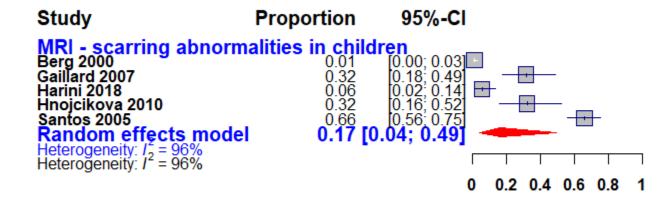


Figure 25: Proportion of scarring abnormalities identified in young people (11 to 25 years old at seizure onset)

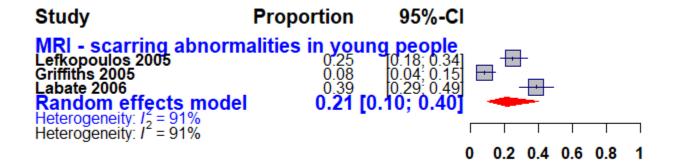


Figure 26: Proportion of scarring abnormalities identified in focal (partial) epilepsy

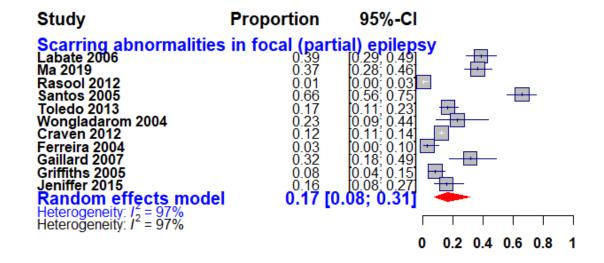


Figure 27: Proportion of scarring abnormalities identified in genetic (idiopathic) generalised epilepsy

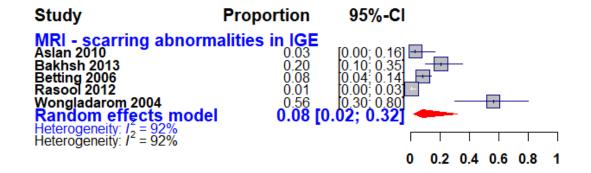


Figure 28: Proportion of scarring abnormalities identified in West syndrome

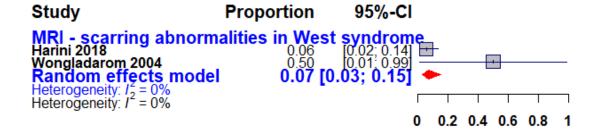


Figure 29: Proportion of scarring abnormalities identified on 1.5-t

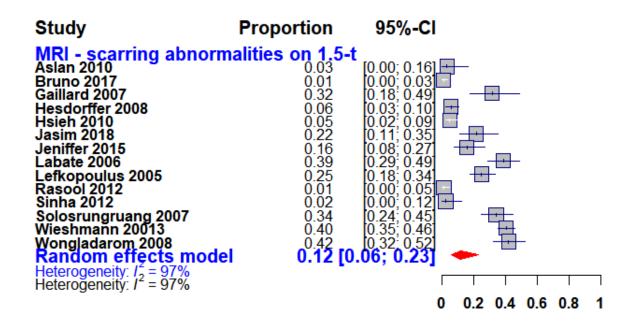


Figure 30: Proportion of scarring abnormalities identified on 3.0-t

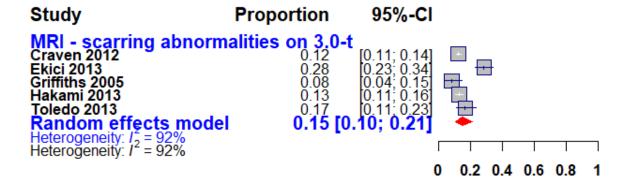


Figure 31: Proportion of scarring abnormalities in those with a new diagnosis

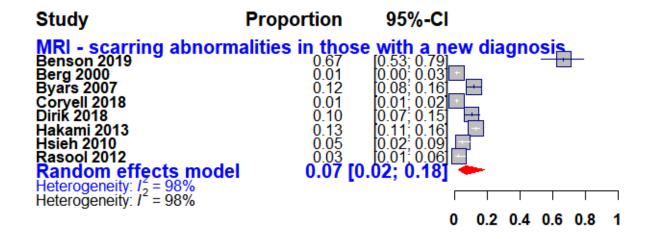


Figure 32: Proportion of scarring abnormalities identified in those with existing diagnosis and treatment resistant

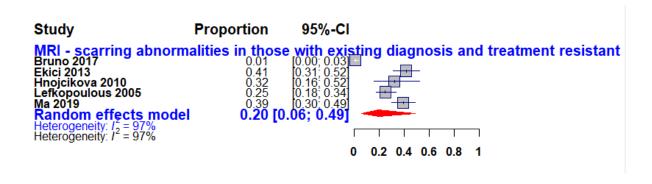


Figure 33: Proportion of scarring abnormalities identified in those with existing diagnosis and controlled epilepsy

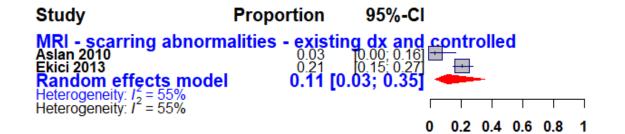


Figure 34: Proportion of scarring abnormalities identified in those without learning disabilities

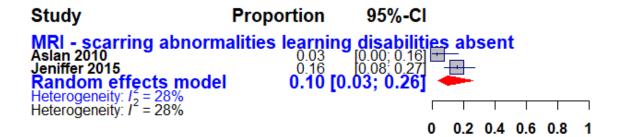
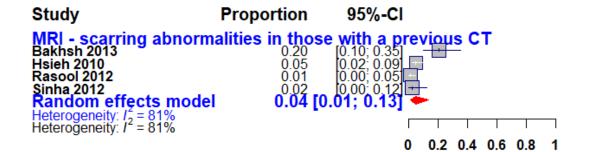


Figure 35: Proportion of scarring abnormalities identified in those with a previous CT scan



Critical outcomes: proportion identified with congenital/developmental abnormalities

Figure 36: Proportion identified with congenital/developmental abnormalities: overall estimate

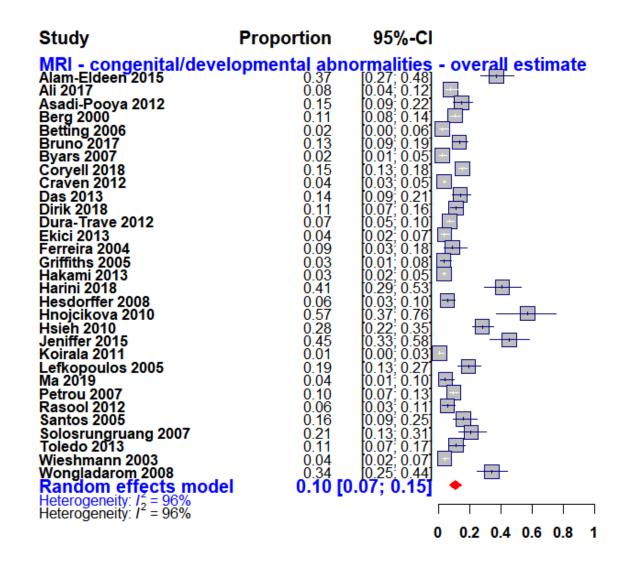


Figure 37: Proportion of congenital/developmental abnormalities indentified in infants (<3 years old at seizure onset)

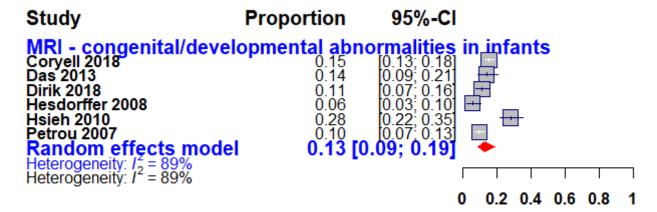


Figure 38: Proportion of congenital/developmental abnormalities identified in children (3 to 11 years old at seizure onset)

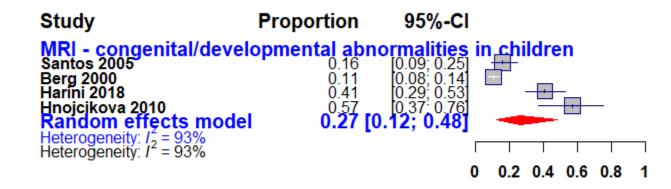


Figure 39: Proportion of congenital/developmental abnormalities identified in young people (11 to 25 years old at seizure onset)

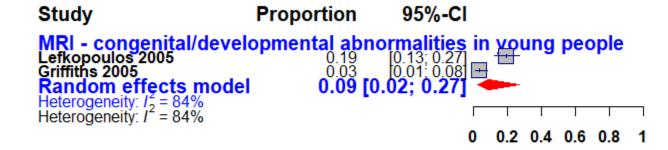


Figure 40: Proportion of congenital/developmental abnormalities identified in focal (partial) epilepsy

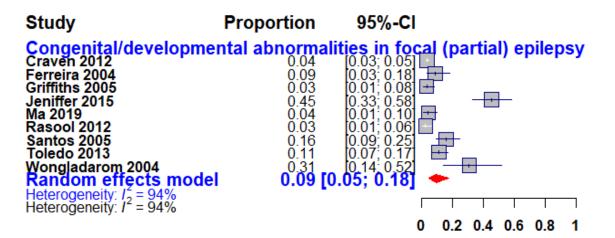


Figure 41: Proportion of congenital/developmental abnormalities identified in genetic (idiopathic) generalised epilepsy

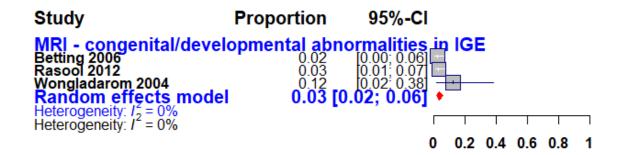


Figure 42: Proportion of congenital/developmental abnormalities identified in West syndrome

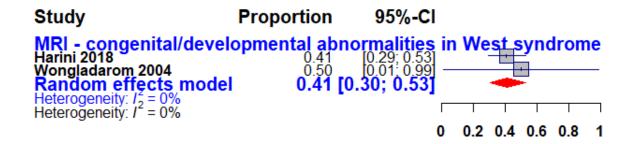


Figure 43: Proportion of congenital/developmental abnormalities identified in Lennox-Gastaut syndrome

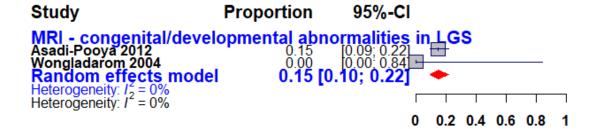


Figure 44: Proportion of congenital/developmental abnormalities identified on 1.5-t

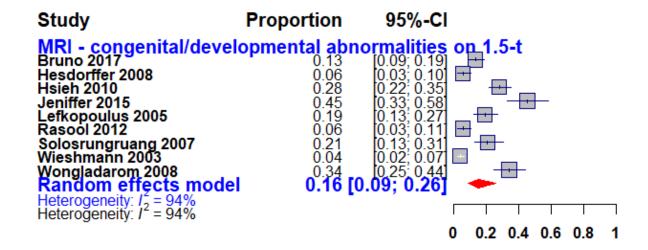


Figure 45: Proportion of congenital/developmental abnormalities identified on 3.0-t

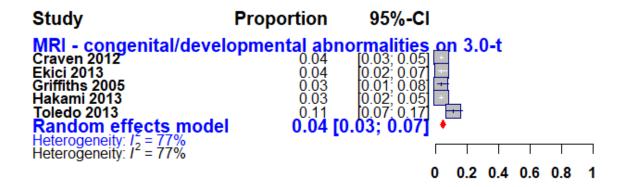


Figure 46: Proportion of congenital/developmental abnormalities identified in those with a new diagnosis

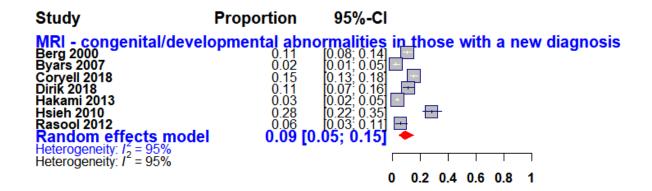


Figure 47: Proportion of congenital/developmental abnormalities identified in those with existing diagnosis and treatment resistant

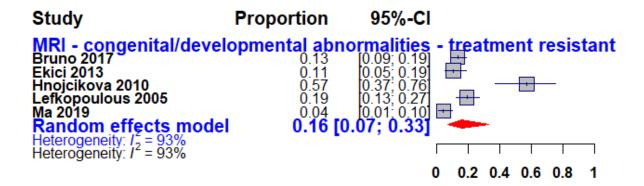
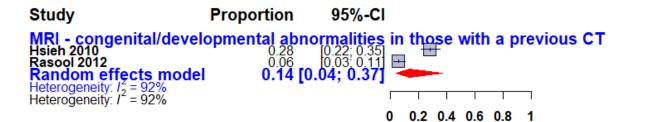


Figure 48: Proportion of congenital/developmental abnormalities identified in those with a previous CT scan



Critical outcomes: proportion identified with inflammatory/infective/immune abnormalities

Figure 49: Proportion identified with inflammatory/infective/immune abnormalities: overall estimate

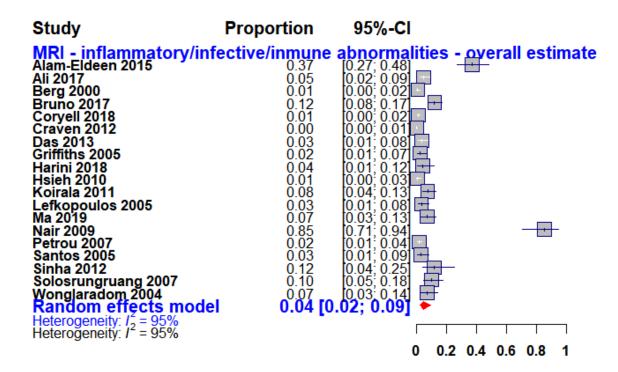


Figure 50: Proportion of inflammatory/infective/inmune abnormalities identified in infants (<3 years old at seizure onset)

Study	Proportion	95%-CI	
MRI - inflammatory/ in Coryell 2018 Das 2013 Hsieh 2010 Petrou 2007 Random effects mode Heterogeneity: I ² = 0% Heterogeneity: I ² = 0%	0.03 0.01	[0.01; 0.08] [0.00; 0.03] [0.01; 0.04] [.01; 0.02]	0.4 0.6 0.8 1

Figure 51: Proportion of inflammatory/infective/inmune abnormalities identified in children (3 to 11 years old at seizure onset)

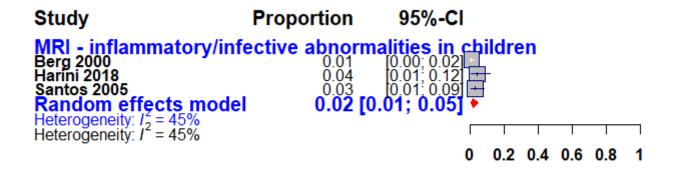


Figure 52: Proportion of inflammatory/infective/inmune abnormalities identified in young people (11 to 25 years old at seizure onset)

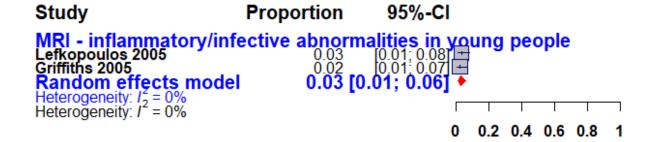


Figure 53: Proportion of inflammatory/infective/inmune abnormalities identified in focal (partial) epilepsy

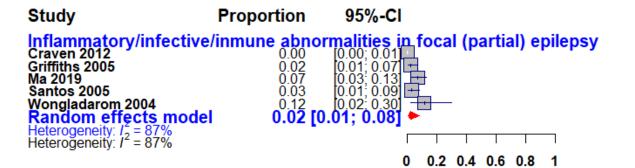


Figure 54: Proportion of inflammatory/infective/inmune abnormalities identified in West syndrome

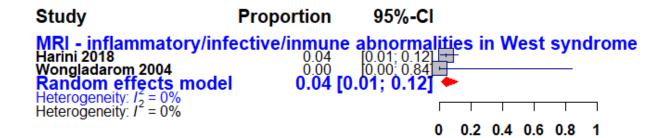


Figure 55: Proportion of inflammatory/infective/inmune abnormalities identified on 1.5-

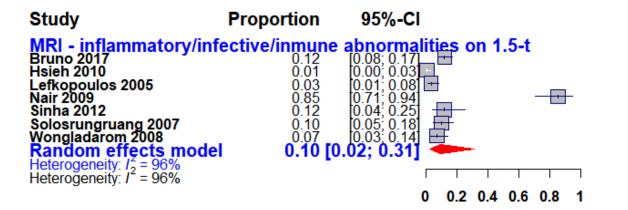


Figure 56: Proportion of inflammatory/infective/inmune abnormalities identified on 3.0-t

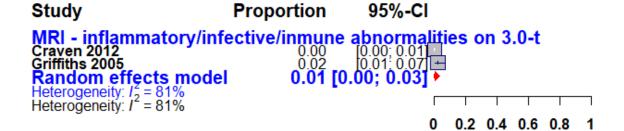


Figure 57:Proportion of inflammatory/infective/inmune abnormalities identified in those with a new diagnosis

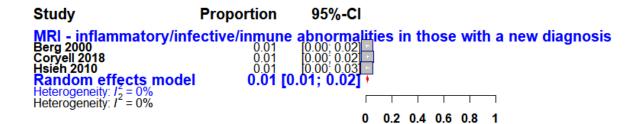


Figure 58: Proportion of inflammatory/infective/inmune abnormalities identified in those with existing diagnosis and treatment resistant

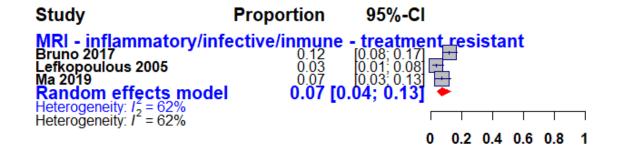
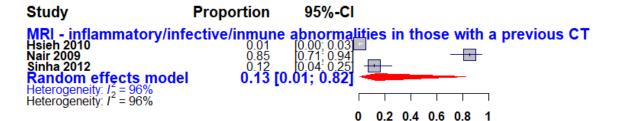


Figure 59: Proportion of inflammatory/infective/inmune abnormalities identified in those with a previous CT scan



Critical outcomes: proportion identified with metabolic/genetic abnormalities

Figure 60: Proportion identified with metabolic/genetic abnormalities: overall estimate

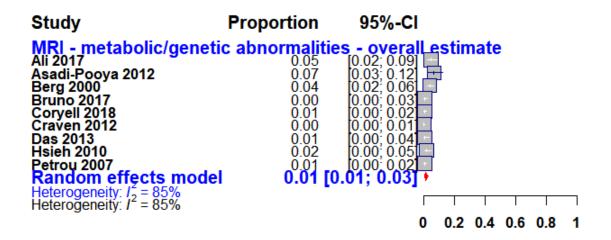


Figure 61: Proportion of metabolic/genetic abnormalities identified in infants (<3 years old at seizure onset)

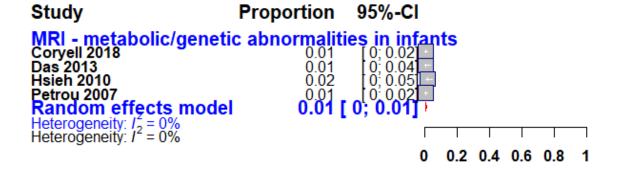


Figure 62: Proportion of metabolic/genetic abnormalities identified on 1.5-t

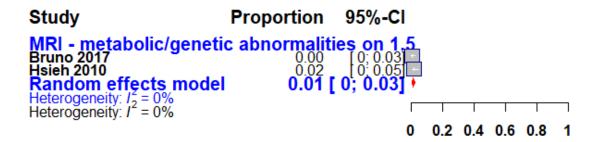
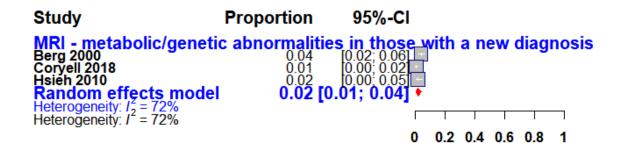


Figure 63: Proportion of metabolic/genetic abnormalities in those with a new diagnosis



Important outcomes: proportion identified with a non-epilepsy related abnormality

Figure 64: Proportion identified with non-epilepsy abnormalities: overall estimate

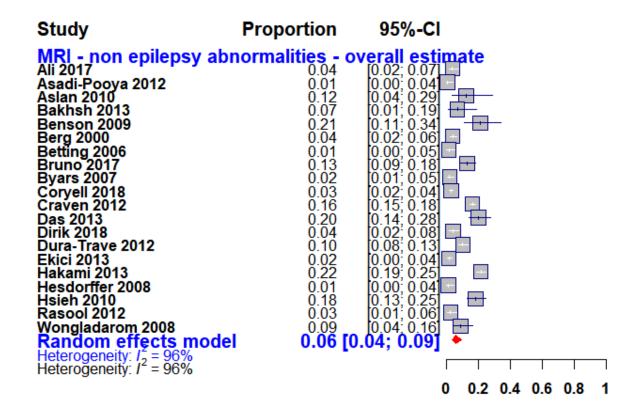


Figure 65: Proportion of non-epilepsy related abnormalities identified in infants (<3 years old at seizure onset)

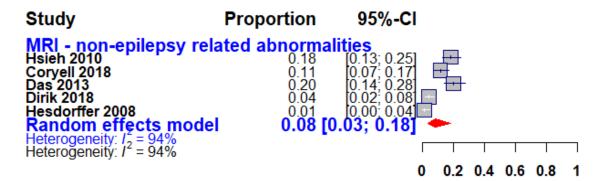


Figure 66: Proportion of non-epilepsy related abnormalities identified in focal (partial) epilepsy

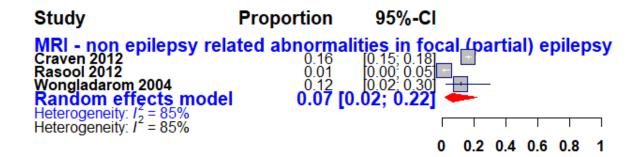


Figure 67: Proportion of non-epilepsy related abnormalities identified in genetic (idiopathic) generalised epilepsy

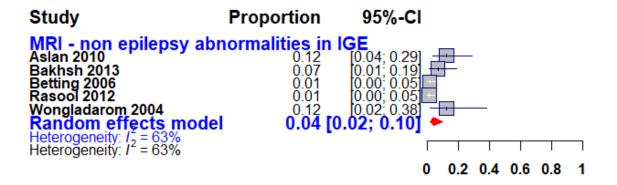


Figure 68: Proportion of non-epilepsy related abnormalities identified in Lennox-Gastaut syndrome

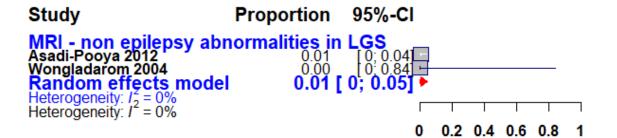


Figure 69: Proportion of non-epilepsy related abnormalities identified on 1.5-t

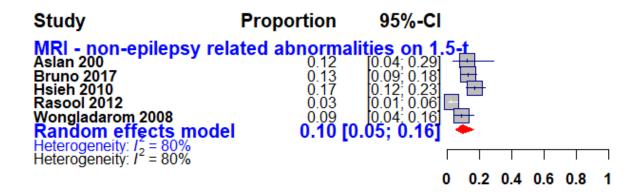


Figure 70: Proportion of non-epilepsy related abnormalities in those with a new diagnosis

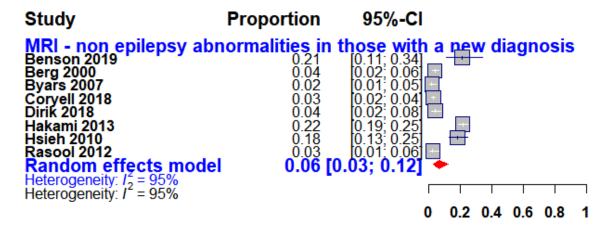


Figure 71: Proportion of non-epilepsy related abnormalities identified in those with an existing diagnosis and treatment resistant

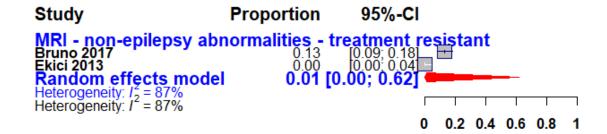


Figure 72: Proportion of non-epilepsy related abnormalities identified in those with an existing diagnosis and controlled

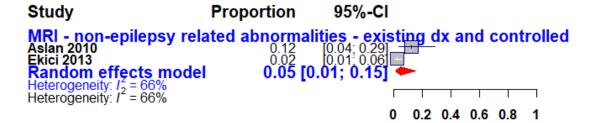
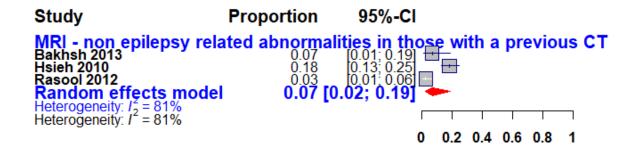


Figure 73: Proportion of non-epilepsy related abnormalities identified in those with a previous CT scan



Appendix F – Adapted GRADE tables

Clinical evidence profile tables for review question: What is the yield of relevant abnormalities detected by MRI in people with epilepsy?

Table 5: Clinical evidence profile for proportion identified with tumour abnormalities

Quality a	Quality assessment					Number of patients				
								Effect		
Number of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Number with a clinically relevant abnormality	Number assessed	Proportion (95% CI)	Quality	Importance
Proportio	on identified with	tumour ab	normalities: ov	verall estimate*						
24 ¹	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Serious ⁴	191	6693	0.03 (0.02 to 0.04)	⊕000 VERY LOW	CRITICAL
Proportio	on of tumour abn	ormalities	identified in inf	ants (<3 years	old at seizure	onset)				
4 ⁵	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁶	11	985	0.01 (0.01 to 0.02)	⊕000 VERY LOW	CRITICAL
Proportio	on of tumour abn	ormalities	identified in ch	ildren (3 to 11 y	ears old at se	eizure onset)				
37	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁶	4	516	0.01 (0 to 0.02)	⊕000 VERY LOW	CRITICAL
Proportio	on of tumour abn	ormalities	identified in yo	ung people (11	to 25 years o	ld at seizure o	nset)			
18	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁶	4	120	0.03 (0.01 to 0.08)	⊕000 VERY LOW	CRITICAL

Quality a	Quality assessment			Number of patients			Effect			
Number of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Number with a clinically relevant abnormality	Number assessed	Proportion (95% CI)	Quality	Importance
Proportio	on of tumour abn	ormalities	identified in old	der people (> 65	years old at	seizure onset)			
1 ⁹	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁶	5	43	0.12 (0.04 to 0.25)	⊕000 VERY LOW	CRITICAL
Proportio	on of tumour abn	ormalities	identified in the	ose with focal (partial) epilep	sy				
7 ¹⁰	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁶	64	2660	0.04 (0.02 to 0.09)	⊕OOO VERY LOW	CRITICAL
Proportio	on of tumour abn	ormalities	identified in the	se with geneti	c (idiopathic)	generalised e	pilepsy			
2 ¹¹	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁶	6	144	0.05 (0.02 to 0.14)	⊕OOO VERY LOW	CRITICAL
Proportio	on of tumour abn	ormalities	identified on 1.	5-t						
8 ¹²	Observational studies	Very serious ²	Serious ¹³	No serious indirectness	Very serious ⁶	49	1080	0.04 (0.02 to 0.07)	⊕OOO VERY LOW	CRITICAL
Proportio	on of tumour abn	ormalities	identified on 3.	0-t						
5 ¹⁴	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Very serious ⁶	71	3309	0.03 (0.01 to 0.06)	⊕000 VERY LOW	CRITICAL
Proportio	on of tumour abn	ormalities	identified in the	ose with a new	diagnosis					
4 ¹⁵	Observational studies	Very serious ²	Serious ¹³	No serious indirectness	Very serious ⁶	31	1556	0.01 (0.00 to 0.03)	⊕OOO VERY LOW	CRITICAL

Quality as	ssessment					Number of p	atients	Effect		
Number of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Number with a clinically relevant abnormality	Number assessed	Proportion (95% CI)	Quality	Importance
Proportio	n of tumour abn	ormalities	identified in the	se with existin	g diagnosis a	nd treatment	resistant			
4 ¹⁶	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Very serious ⁶	27	454	0.05 (0.02 to 0.12)	⊕000 VERY LOW	CRITICAL
Proportio	n of tumour abn	ormalities	identified in the	se with existin	g diagnosis a	ind controlled				
1 ¹⁷	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁶	0	170	0.00 (0 to 0.02)	⊕000 VERY LOW	CRITICAL
Proportio	n of tumour abn	ormalities	identified in the	se without lea	rning disabilit	ies				
1 ¹⁸	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁶	1	64	0.02 (0 to 0.08)	⊕000 VERY LOW	CRITICAL
Proportio	n of tumour abn	ormalities	identified in the	ose who had a	previous CT s	can				
3 ¹⁹	Observational studies	Very serious ²	Serious ¹³	No serious indirectness	Very serious ⁶	10	269	0.04 (0.01 to 0.13)	⊕000 VERY LOW	CRITICAL

¹ Ali 2017, Bakhsh 2013, Berg 2000, Bruno 2017, Craven 2012, Das 2013, Dirik 2018, Dura-Trave 2012, Ekici 2013, Griffiths 2005, Hakami 2013, Hnojcikova 2010, Hsieh 2010, Jasim 2018, Jeniffer 2015, Koirala 2011, Ma 2019, Petrou 2007, Santos 2005, Sinha 2012, Solosrungrouang 2007, Toledo 2013, Wieshmann 2003, Wongladarom 2004

² Very serious risk of bias in the evidence contributing to the outcomes as per CEBMA checklist

³ Very serious heterogeneity (*I*²>75%)

⁴ Number of events >150 but <300

⁵ Das 2013, Diriki 2018, Hsieh 2010, Petrou 2007

⁶ Number of events <150

⁷ Berg 2000, Hnojcikova 2010, Santos 2005

⁸ Griffiths 2005

⁹ Sinha 2012

18 Jenniffer 2015

Table 6: Clinical evidence profile for proportion identified with vascular abnormalities

Quality as	ssessment					Number of patients		Effect		
Number of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Number with a clinically relevant abnormality	Number assessed	Proportion (95% CI)	Quality	Importance
Proportio	n identified with	vascular a	abnormalities: d	verall estimate	<u>4</u>					
25 ¹	Observational studies	Very serious ²	Very serious ³	No serious indirectness	No serious imprecision	412	7544	0.06 (0.04 to 0.8)	⊕000 VERY LOW	CRITICAL
Proportio	n of vascular ab	normalitie	s identified in c	hildren (3 to 11	years old at	seizure onset)				
34	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Very serious ⁵	27	559	0.04 (0.01 to 0.18)	⊕000 VERY LOW	CRITICAL
Proportio	n of vascular ab	normalitie	s identified in y	oung people (1	1 to 25 years	old at seizure	onset)			
26	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁵	16	240	0.07 (0.04 to 0.48)	⊕000 VERY LOW	CRITICAL
Proportio	n of vascular ab	normalitie	s identified in o	lder people (> 6	65 years old a	t seizure onse	et)			

¹⁰ Craven 2012, Griffiths 2005, Jeniffer 2015, Ma 2019, Santos 2005, Toledo 2013, Wongladarom 2004

¹¹ Bakhsh 2012, Wongladarom 2004

¹² Bruno 2017, Hsieh 2010, Jasim 2018, Jeniffer 2015, Sinha 2012, Solosrungruang 2007, Wieshmann 2013, Wongladarom 2004

¹³ Serious heterogeneity (*I*² >50% but <75%)

¹⁴ Craven 2012, Ekici 2013, Griffiths 2005, Hakami 2013, Toledo 2013

¹⁵ Berg 2000, Dirik 2018, Hakami 2013, Hsieh 2010

¹⁶ Bruno 2017, Ekici 2013, Hnojcikova 2010, Ma 2019

¹⁷ Ekici 2013

¹⁹ Bakhsh 2013, Hsieh 2010, Sinha 2012

Quality as	ssessment					Number of p	lumber of patients Effect			
Number of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Number with a clinically relevant abnormality	Number assessed	Proportion (95% CI)	Quality	Importance
1 ⁷	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁵	13	43	0.30 (0.17 to 0.46)	⊕000 VERY LOW	CRITICAL
Proportio	on of vascular ab	normalities	s identified in th	nose with focal	(partial) epile	psy				
68	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Very serious ⁵	68	2596	0.04 (0.02 to 0.08)	⊕OOO VERY LOW	CRITICAL
Proportio	on of vascular ab	normalities	s identified in th	nose with gene	tic (idiopathic) generalised	epilepsy			
2 ⁹	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁵	5	60	0.08 (0.04 to 0.19)	⊕OOO VERY LOW	CRITICAL
Proportio	on of vascular ab	normalities	s identified in th	nose with West	syndrome					
2 ¹⁰	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁵	15	73	0.21 (0.13 to 0.31)	⊕OOO VERY LOW	CRITICAL
Proportio	on of vascular ab	normalities	s identified in th	nose with Lenn	ox-Gastaut sy	ndrome				
111	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁵	0	1	0.00 (0 to 0.02)	⊕OOO VERY LOW	CRITICAL
Proporti	ion of vascular a	bnormalitie	es identified on	1.5-t						
7 ¹²	Observational studies	Very serious ²	Serious ¹³	No serious indirectness	Very serious ⁵	85	794	0.11 (0.07 to 0.17)	⊕OOO VERY LOW	CRITICAL
Proportio	on of vascular ab	normalities	s identified on 3	3.0-t						

Quality a	ssessment					Number of p	atients	Effect		
Number of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Number with a clinically relevant abnormality	Number assessed	Proportion (95% CI)	Quality	Importance
5 ¹⁴	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Very serious ⁵	27	559	0.04 (0.02 to 0.07)	⊕OOO VERY LOW	CRITICAL
Proportio	on of vascular ab	normalities	s identified in th	nose with a nev	v diagnosis [△]					
615	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Serious ¹⁶	119	2370	0.04 (0.02 to 0.09)	⊕000 VERY LOW	CRITICAL
Proportio	on of vascular ab	normalities	s identified in th	nose with exist	ing diagnosis	and treatmen	t resistar	nt		
3 ¹⁷	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁵	27	426	0.06 (0.04 to 0.09)	⊕OOO VERY LOW	CRITICAL
Proportio	on of vascular ab	normalities	s identified in th	nose with exist	ing diagnosis	and controlle	d			
1 ¹⁸	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁵	3	170	0.02 (0 to 0.05)	⊕000 VERY LOW	CRITICAL

Δ One of the included studies (Benson 2019) included people with arteriovenous malformations (AVM) only, which may overestimate the yield of identified vascular abnormalities 1 Alam-Eldeen 2015, Ali 2017, Bakhsh 2013, Berg 2000, Bruno 2017, Coryell 2018, Craven 2012, Das 2013, Dirik 2018, Dura-Trave 2012, Ekici 2013, Griffiths 2005, Hakami 2013, Harini 2018, Hsieh 2010, Koirala 2011, Lefkopoulos 2005, Ma 2019, Nair 2009, Petrou 2007, Santos 2005, Solosrungrouang 2007, Toledo 2013, Wieshmann 2003, Wongladarom 2004

- 2 Very serious risk of bias in the evidence contributing to the outcomes as per CEBMA checklist
- 3 Very serious heterogeneity (I²>75%)
- 4 Berg 2000, Harini 2018, Santos 2005
- 5 Number of events <150
- 6 Griffiths 2005, Lefkopoulos 2005
- 7 Sinha 2012
- 8 Craven 2012, Griffiths 2005, Ma 2019, Santos 2005, Toledo 2013, Wongladarom 2004
- 9 Bakhsh 2013, Wongladarom 2004

Table 7: Clinical evidence profile for proportion identified with scarring abnormalities

Quality as	ssessment					Number of p	patients	Effect		
Number of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Number with a clinically relevant abnormality	Number assessed	Proportion (95% CI)	Quality	Importance
Proportio	n identified with	scarring a	bnormalities: o	verall estimate						
37 ¹	Observational studies	Very serious ²	Very serious ³	No serious indirectness	No serious imprecision	1146	8681	0.10 (0.06 to 0.16)	⊕000 VERY LOW	CRITICAL
Proportio	n of scarring abi	normalities	identified in in	fants (<3 vears	old at seizur	e onset)				
6 ⁴	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Very serious ⁵	73	1858	0.04 (0.02 to 0.09)	⊕000 VERY LOW	CRITICAL
Proportio	n of scarring abı	normalities	identified in cl	hildren (3 to 11	years old at s	seizure onset)				

¹⁰ Harini 2018, Wongladarom 2004

¹¹ Wongladarom 2004

¹² Bruno 2017, Hsieh 2010, Lefkopoukus 2005, Nair 2009, Sinha 2012, Solosrungruang 2007, Wongladarom 2004

¹³ Serious heterogeneity (*I*² >50% but <75%)

¹⁴ Craven 2012, Ekici 2013, Griffiths 2005, Hakami 2013, Toledo 2013

¹⁵ Berg 2000, Coryell 2008, Dirik 2018, Hakami 2013, Hsieh 2010

¹⁶ Number of events >150 but <300

¹⁷ Bruno 2017, Ekici 2013, Ma 2019

¹⁸ Ekici 2013

Quality as	ssessment					Number of p	patients	Effect		
Number of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Number with a clinically relevant abnormality	Number assessed	Proportion (95% CI)	Quality	Importance
5 ⁶	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Very serious ⁵	96	625	0.17 (0.04 to 0.49)	⊕000 VERY LOW	CRITICAL
Proportio	n of scarring ab	normalities	s identified in yo	oung people (1	1 to 25 years	old at seizure	onset)			
37	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Very serious ⁵	79	341	0.21 (0.10 to 0.40)	⊕000 VERY LOW	CRITICAL
Proportio	on of scarring ab	normalities	s identified in a	dults (25 to 65 y	years old sat	seizure onset)			
18	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁵	11	134	0.08 (0.04 to 0.14)	⊕000 VERY LOW	CRITICAL
Proportio	on of scarring ab	normalities	s identified in o	lder people (> 6	55 years old a	t seizure onse	et)			
1 ⁹	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁵	1	43	0.02 (0 to 0.12)	⊕000 VERY LOW	CRITICAL
Proportio	on of scarring ab	normalities	s identified in th	nose with focal	(partial) epile	psy				
11 ¹⁰	Observational studies	Very serious ²	Very serious ³	No serious indirectness	No serious imprecision	500	3023	0.17 (0.08 to 0.31)	⊕000 VERY LOW	CRITICAL

Quality as	ssessment					Number of p	atients	Effect		
Number of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Number with a clinically relevant abnormality	Number assessed	Proportion (95% CI)	Quality	Importance
Proportio	n of scarring ab	normalities	s identified in th	nose with genet	tic (idiopathic) generalised	epilepsy			
5 ¹¹	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Very serious ⁵	65	467	0.08 (0.02 to 0.32)	⊕000 VERY LOW	CRITICAL
Proportio	n of scarring abi	normalities	in those with \	Nest syndrome) }					
2 ¹²	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁵	46	171	0.07 (0.03 to 0.15)	⊕000 VERY LOW	CRITICAL
Proportio	n of scarring abi	normalities	s identified in th	ose with Lenn	ox-Gastaut sy	ndrome				
1 ¹³	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁵	42	100	0.42 (0.32 to 0.52)	⊕000 VERY LOW	CRITICAL
Proportio	n of scarring abi	normalities	identified on 1	.5-t						
14 ¹⁴	Observational studies	Very serious ²	Very serious ³	No serious indirectness	No serious imprecision	331	1687	0.12 (0.06 to 0.23)	⊕000 VERY LOW	CRITICAL
Proportio	n of scarring ab	normalities	s identified on 3	3.0-t						

Quality as	ssessment					Number of p	patients	Effect		
Number of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Number with a clinically relevant abnormality	Number assessed	Proportion (95% CI)	Quality	Importance
5 ¹⁵	Observational studies	Very serious ²	Very serious ³	No serious indirectness	No serious imprecision	458	3045	0.15 (0.10 to 0.21)	⊕000 VERY LOW	CRITICAL
Proportio	n of scarring ab	normalities	s identified in th	ose with a new	diagnosis					
8 ¹⁶	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Serious ¹⁷	212	2576	0.07 (0.02 to 0.18)	⊕000 VERY LOW	CRITICAL
Proportio	on of scarring ab	normalities	s identified in th	nose with existi	ng diagnosis	and treatmen	t resistar	nt		
5 ¹⁸	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Very serious ⁵	122	574	0.20 (0.06 to 0.49)	⊕000 VERY LOW	CRITICAL
Proportio	on of scarring ab	normalities	s identified in th	ose with existi	ng diagnosis	and controlle	d			
2 ¹⁹	Observational studies	Very serious ²	Serious ²⁰	No serious indirectness	Very serious ⁵	36	202	0.11 (0.03 to 0.35)	⊕000 VERY LOW	CRITICAL
Proportio	on of scarring ab	normalities	s identified in th	nose without lea	arning disabil	ities				
2 ²¹	Observational studies	Very serious ²	No serious inconsistency	Serious ²²	Very serious ⁵	11	96	0.10 (0.03 to 0.26)	⊕000 VERY LOW	CRITICAL

Quality as	Quality assessment					Number of p	oatients	Effect		
Number of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Number with a clinically relevant abnormality	Number assessed	Proportion (95% CI)	Quality	Importance
Proportio	on of scarring ab	normalities	s identified in th	nose who had a	previous C1	scan				
4 ²³	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Very serious ⁵	21	426	0.04 (0.01 to 0.13)	⊕000 VERY LOW	CRITICAL

¹ Alam-Eldeen 2015, Ali 2017, Aslan 2010, Bakhsh 2013, Benson 2009, Berg 2000, Betting 2006, Bruno 2017, Byars 2007, Coryell 2018, Craven 2012, Das 2013, Dirik 2018, Dura-Trave 2012, Ekici 2013, Ferreira 2004, Gaillard 2007, Griffiths 2005, Hakami 2013, Harini 2018, Hersdorffer 2008, Hnojcikova 2010, Hsieh 2010, Jeniffer 2015, Jasim 2018, Koirala 2011, Labate 2006, Lefkopoulos 2005, Ma 2019, Petrou 2007, Rasool 2012, Santos 2005, Sinha 2012, Solosrungruang 2007, Toledo 2013, Wieshmann 2003, Wongladarom 2004

- 2 Very serious risk of bias in the evidence contributing to the outcomes as per CEBMA checklist
- 3 Very serious heterogeneity (I²>75%)
- 4 Coryell 2018, Das 2013, Dirik 2018, Hesdorffer 2008, Hsieh 2010, Petrou 2007
- 5 Number of events <150
- 6 Berg 2000, Gaillard 2007, Harini 2018, Hnojcikova 2010, Santos 2005
- 7 Lefkopoulos 2005, Griffiths 2005, Labate 2006
- 8 Betting 2006
- 9 Sinha 2012
- 10 Craven 2012, Ferreira 2004, Gaillard 2007, Griffiths 2005, Jeniffer 2015, Labate 2006, Ma 2019, Rasool 2012, Santos 2005, Toledo 2013, Wongladarom 2004
- 11 Aslan 2010, Bakhsh 2013, Betting 2006, Rasool 2012, Wongladarom 2004
- 12 Harini 2018, Wongladarom 2004
- 13 Wongladarom 2004
- 14 Aslan 2010, Bruno 2017, Gaillard 2007, Hesdorffer 2008, Hsieh 2010, Jasim 2018, Jeniffer 2015, Labate 2006, Lefkopoulos 2005, Rasool 2012, Sinha 2012, Solosgruang 2007, Wieshmann 2013, Wongladarom 2004
- 15 Craven 2012, Ekici 2013, Griffiths 2005, Hakami 2013, Toledo 2013
- 16 Benson 2019, Berg 2000, Byars 2007, Coryell 2018, Dirik 2018, Hakami 2013, Hsieh 2010, Rasool 2012
- 17 Number of events >150 but <300
- 18 Bruno 2017, Ekici 2013, Hnojcikova 2010, Lefkopoulos 2005, Ma 2019

Table 8: Clinical evidence profile for proportion identified with congenital/developmental abnormalities

Quality as	ssessment					Number of p	atients	Effect		
Number of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Number with a clinically relevant abnormality	Number assessed	Proportion (95% CI)	Quality	Importance
Proportio	on identidied with	n congenita	al/development	al abnormalitie	s: overall esti	mate				
31 ¹	Observational studies	Very serious ²	Very serious ³	No serious indirectness	No serious imprecision	768	8450	0.10 (0.07 to 0.15)	⊕000 VERY LOW	CRITICAL
Proportio	on of congenital/o	developme	ntal abnormalit	ies identified ir	n infants (<3 y	ears old at se	izure ons	set)		
64	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Serious ⁵	256	1858	0.13 (0.09 to 0.19)	⊕000 VERY LOW	CRITICAL
Proportio	on of congenital/o	developme	ntal abnormalit	ies identified ir	n children (3 t	o 11 years old	at seizu	re onset)		
4 ⁶	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Very serious ⁷	102	587	0.27 (0.12 to 0.48)	⊕000 VERY LOW	CRITICAL

¹⁹ Aslan 2010, Ekici 2013

²⁰ Serious heterogeneity (I² >50% but <75%)

²¹ Aslan 2010, Jenifer 2015

²² Population is indirect in 1 study (3% of participants did have learning disabilities) 23 Bakhsh 2013, Hsieh 2010, Rasool 2012, Sinha 2012

Quality as	ssessment					Number of p	atients	Effect		
Number of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Number with a clinically relevant abnormality	Number assessed	Proportion (95% CI)	Quality	Importance
Proportio	n of congenital/o	developme	ntal abnormalit	ies identified ir	n young peop	le (11 to 25 ye	ars old a	t seizure onse	et)	
28	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Very serious ⁷	27	240	0.09 (0.02 to 0.27)	⊕000 VERY LOW	CRITICAL
Proportio	n of congenital/o	developme	ntal abnormalit	ies identified ir	n adults (25 to	65 years old	at seizur	e onset)		
1 ⁹	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁷	3	134	0.02 (0 to 0.06)	⊕000 VERY LOW	CRITICAL
Proportio	n of congenital/o	developme	ntal abnormalit	ies identified ir	n those with f	ocal (partial) e	pilepsy			
910	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Serious ⁵	168	2810	0.09 (0.05 to 0.18)	⊕000 VERY LOW	CRITICAL
Proportio	n of congenital/o	developme	ntal abnormalit	ies identified in	n those with g	jenetic (idiopa	ithic) gen	eralised epile	psy	
3 ¹¹	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁷	14	307	0.03 (0.02 to 0.06)	⊕000 VERY LOW	CRITICAL
Proportio	n of congenital/o	developme	ntal abnormalit	ies identified ir	n those with V	Vest syndrom	e			

Quality as	ssessment					Number of p	atients	Effect		
Number of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Number with a clinically relevant abnormality	Number assessed	Proportion (95% CI)	Quality	Importance
212	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁷	30	73	0.41 (0.30 to 0.53)	⊕000 VERY LOW	CRITICAL
Proportio	on of congenital/	developme	ntal abnormalit	ies identified ir	n those with L	.ennox-Gastaı	ut syndro	me		
2 ¹³	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁷	20	137	0.15 (0.10 to 0.22)	⊕000 VERY LOW	CRITICAL
Proportio	on of congenital/	developme	ntal abnormalit	ies identified o	n 1.5-t					
914	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Serious ⁵	216	1422	0.16 (0.09 to 0.26)	⊕000 VERY LOW	CRITICAL
Proportio	on of congenital/	developme	ntal abnormalit	ies identified o	n 3.0-t					
5 ¹⁵	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Very serious ⁷	131	3309	0.04 (0.03 to 0.07)	⊕000 VERY LOW	CRITICAL
Proportio	on of congenital/	developme	ntal abnormalit	ies identified ir	n those with a	new diagnos	is			

Quality as	ssessment					Number of p	atients	Effect		
Number of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Number with a clinically relevant abnormality	Number assessed	Proportion (95% CI)	Quality	Importance
7 ¹⁶	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Serious ⁵	267	2676	0.09 (0.05 to 0.15)	⊕000 VERY LOW	CRITICAL
Proportio	on of congenital/o	developme	ntal abnormalit	ies identified ir	n those with e	xisting diagno	osis and	treatment resi	stant	
5 ¹⁷	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Very serious ⁷	83	574	0.16 (0.07 to 0.33)	⊕000 VERY LOW	CRITICAL
Proportio	on of congenital/o	developme	ntal abnormalit	ies identified ir	n those with e	xisting diagno	osis and	controlled		
1 ¹⁸	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁷	0	170	0.00 (0 to 0.02)	⊕000 VERY LOW	CRITICAL
Proportio	on of congenital/o	developme	ntal abnormalit	ies identified ir	n those with le	earning disabi	ilities			
1 ¹⁹	Observational studies	Very serious ²	No serious inconsistency	Serious ²⁰	Very serious ⁷	20	135	0.15 (0.09 to 0.22)	⊕000 VERY LOW	CRITICAL
Proportio	oportion of congenital/developmental abnormalities identified					ut learning dis	sabilities			

Quality as	ssessment					Number of p	atients	Effect		
Number of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Number with a clinically relevant abnormality	Number assessed	Proportion (95% CI)	Quality	Importance
1 ²¹	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁷	29	64	0.45 (0.33 to 0.58)	⊕000 VERY LOW	CRITICAL
Proportio	on of congenital/o	developme	ntal abnormalit	ies identified ir	n those who h	ad a previous	CT scan			
2 ²²	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Very serious ⁷	60	339	0.14 (0.04 to 0.37)	⊕000 VERY LOW	CRITICAL

¹ Alam-Eldeen 2015, Ali 2017, Asadi-Pooya 2012, Berg 2000, Betting 2006, Bruno 2017, Byars 2007, Coryell 2018, Craven 2012, Das 2013, Dirik 2018, Dura-Trave 2012, Ekici 2013, Ferreira 2004, Griffiths 2005, Hakami 2013, Harini 2018, Hersdorffer 2008, Hnojcikova 2010, Hsieh 2010, Jeniffer 2015, Koirala 2011, Lefkopoulos 2005, Ma 2019, Petrou 2007, Rasool 2012, Santos 2005, Solosrungruang 2007, Toledo 2013, Wieshmann 2003, Wongladarom 2004

- 2 Very serious risk of bias in the evidence contributing to the outcomes as per CEBMA checklist
- 3 Very serious heterogeneity (I²>75%)
- 4 Coryell 2018, Das 2013, Dirik 2018, Hesdorffer 2008, Hsieh 2010, Petrou 2007
- 5 Number of events >150 but <300
- 6 Santos 2005, Berg 2000, Harini 2018, Hnojcikova 2010
- 7 Number of events <150
- 8 Lefkopoulos 2005, Griffiths 2005
- 9 Betting 2006
- 10 Craven 2012, Ferreira 2004, Griffiths 2005, Jeniffer 2015, Ma 2019, Rasool 2012, Santos 2005, Toledo 2013, Wongladarom 2004
- 11 Betting 2006, Rasool 2012, Wongladarom 2004
- 12 Harini 2018, Wongladarom 2004
- 13 Asadi-Pooya 2012, Wongladarom 2004
- 14 Bruno 2017, Hesdorffer 2008, Hsieh 2010, Jeniffer 2015, Lefkopoulos 2005, Rasool 2012, Solosgruang 2007, Wieshmann 2013, Wongladarom 2004

15 Craven 2012, Ekici 2013, Griffiths 2005, Hakami 2013, Toledo 2013
16 Berg 2000, Byars 2007, Coryell 2018, Dirik 2018, Hakami 2013, Hsieh 2010, Rasool 2012
17 Bruno 2017, Ekici 2013, Hnojcikova 2010, Lefkopoulos 2005, Ma 2019
18 Ekici 2013
19 Asadi-Pooya 2012
20 Population is indirect in 1 study (3% of participants did not have learning disabilities)
21 Jeniffer 2015
22 Hsieh 2010, Rasool 2012

Table 9: Clinical evidence profile for proportion identified with inflammatory/infective/immune abnormalities

Quality a	ssessment		, ,		,	Number of p	atients	Effect		
Number of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Number with a clinically relevant abnormality	Number assessed	Proportion (95% CI)	Quality	Importance
Proportio	n identified with	inflammat	ory/infective/im	nmune abnorma	alities: overall	estimate [¥]				
19 ¹	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Serious ⁴	187	5341	0.04 (0.02 to 0.09)	⊕000 VERY LOW	CRITICAL
Proportio	on of inflammator	ry/infective	e/immune abnoi	rmalities identif	ied in infants	(<3 years old	at seizur	e onset)		
4 ⁵	Observational Very No serious No serious inconsistency indirectness					22	1477	0.01 (0.01 to 0.02)	⊕000 VERY LOW	CRITICAL
Proportio	Proportion of inflammatory/infective/immune abnormalities ident					n (3 to 11 yea	rs old at	seizure onset		

Quality a	Quality assessment					Number of patients Effect				
Number of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Number with a clinically relevant abnormality	Number assessed	Proportion (95% CI)	Quality	Importance
37	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁶	9	559	0.02 (0.01 to 0.05)	⊕000 VERY LOW	CRITICAL
Proportio	on of inflammato	ry/infective	/immune abnoi	malities identif	fied in young	people (11 to	25 years	old at seizure	onset)	
28	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁶	7	240	0.03 (0.01 to 0.06)	⊕000 VERY LOW	CRITICAL
Proportio	on of inflammator	ry/infective	e/immune abnoi	malities identif	fied in older p	eople (> 65 ye	ears old a	t seizure onse	et)	
1 ⁹	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁶	5	43	0.12 (0.04 to 0.25)	⊕000 VERY LOW	CRITICAL
Proportio	on of inflammator	ry/infective	e/immune abnoi	malities identif	fied in those v	vith focal (par	tial) epile	psy		
5 ¹⁰	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Very serious ⁶	21	2361	0.02 (0.01 to 0.08)	⊕000 VERY LOW	CRITICAL
Proportio	on of inflammato	ry/infective	/immune abnoi	malities identif	fied in those v	vith genetic (i	diopathic) generalised	epilepsy	
1 ¹¹	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁶	2	16	0.12 (0.02 to 0.38)	⊕000 VERY LOW	CRITICAL

Quality as	uality assessment				Number of patients Effect			Effect		
Number of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Number with a clinically relevant abnormality	Number assessed	Proportion (95% CI)	Quality	Importance
Proportio	n of inflammator	ry/infective	/immune abnoi	malities identif	fied in those v	vith West syn	drome			
212	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁶	3	73	0.04 (0.01 to 0.12)	⊕000 VERY LOW	CRITICAL
Proportio	n of inflammator	ry/infective	/immune abnoi	malities identif	fied in those v	vith Lennox-G	astaut sy	/ndrome		
111	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁶	0	2	0.00 (0 to 0.02)	⊕000 VERY LOW	CRITICAL
Proportio	n of inflammator	ry/infective	/immune abnoi	malities identif	fied on 1.5-t [¥]					
7 ¹³	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Very serious ⁶	87	794	0.10 (0.02 to 0.31)	⊕000 VERY LOW	CRITICAL
Proportio	n of inflammator	ry/infective	/immune abnoi	malities identif	fied on 3.0-t					
214	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Very serious ⁶	7	2120	0.01 (0.00 to 0.03)	⊕000 VERY LOW	CRITICAL
Proportio	n of inflammator	y/infective	/immune abnoi	malities in thos	se with a new	diagnosis				

Quality a	Quality assessment					Number of p	atients	Effect		
Number of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Number with a clinically relevant abnormality	Number assessed	Proportion (95% CI)	Quality	Importance
3 ¹⁵	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁶	12	1284	0.01 (0.01 to 0.02)	⊕000 VERY LOW	CRITICAL
Proportio	on of inflammato	ry/infective	/ /immune abnoi	malities identif	fied in those v	vith existing d	liagnosis	and treatmen	t resistant	¥
3 ¹⁶	Observational studies	Very serious ²	Serious ¹⁷	No serious indirectness	Very serious ⁶	38	452	0.07 (0.04 to 0.13)	⊕000 VERY LOW	CRITICAL
Proportio	on of inflammato	ry/infective	/immune abnoi	malities identif	fied in those v	vho had a pre	vious CT	scan		
318	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Very serious ⁶	41	266	0.13 (0.01 to 0.82)	⊕000 VERY LOW	CRITICAL

[¥] In 1 of the included studies (Bruno 2017), all infections identified were neurocysticercosis, which is a condition endemic to Bhutan, where the study was conducted 1 Alam-Eldeen 2015, Ali 2017, Berg 2000, Bruno 2017, Coryell 2018, Craven 2012, Das 2013, Griffiths 2005, Harini 2018, Hsieh 2010, Koirala 2011, Lefkopoulos 2005, Ma 2019, Nair 2009, Petrou 2007, Santos 2005, Sinha 2012, Solosrungruang 2007, Wongladarom 2004

² Very serious risk of bias in the evidence contributing to the outcomes as per CEBMA checklist

³ Very serious heterogeneity (I²>75%)

⁴ Number of events >150 but <300

⁵ Coryell 2018, Das 2013, Hsieh 2010, Petrou 2007

⁶ Number of events <150

⁷ Berg 2000, Harini 2018, Santos 2005

⁸ Lefkopoulos 2005, Griffiths 2005

⁹ Sinha 2012

Table 10: Clinical evidence profile for proportion identified with metabolic/genetic abnormalities

	ssessment					Number of p	atients			
								Effect		
Number of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Number with a clinically relevant abnormality	Number assessed	Proportion (95% CI)	Quality	Importance
Proportio	on identified with	metabolic	/genetic abnorr	nalities: overal	estimate					
91	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Very serious ⁴	54	4426	0.01 (0.01 to 0.03)	⊕000 VERY LOW	CRITICAL
Proportio	on of metabolic/g	enetic abn	ormalities iden	tified in infants	(<3 vears old	at seizure on	set)			
4 ⁵	Observational studies	Very serious ₂	No serious inconsistency	No serious indirectness	Very serious ⁴	10	1477	0.01 (0 to 0.01)	⊕000 VERY LOW	CRITICAL
Proportio	on of metabolic/g	enetic abn	ormalities iden	tified in childre	n (3 to 11 yea	rs old at seizu	ıre onset)		

¹⁰ Craven 2012, Griffiths 2005, Ma 2019, Santos 2005, Wongladarom 2004

¹¹ Wongladarom 2004

¹² Harini 2018, Wongladarom 2004

¹³ Bruno 2017, Hsieh 2010, Lefkopoulos 2005, Nair 2009, Sinha 2012, Solosrungruang 2007, Wongladarom 2004

¹⁴ Craven 2012, Griffiths 2005

¹⁵ Berg 2000, Coryell 2018, Hsieh 2010

¹⁶ Bruno 2017, Lefkopoulos 2005, Ma 2019

¹⁷ Serious heterogeneity (I² >50% but <75%)

¹⁸ Hsieh 2010, Nair 2009, Sinha 2012

Quality as	Quality assessment				Number of p	Effect				
Number of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Number with a clinically relevant abnormality	Number assessed	Proportion (95% CI)	Quality	Importance
1 ⁶	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁴	15	388	0.04 (0.02 to 0.06)	⊕000 VERY LOW	CRITICAL
Proportio	n of metabolic/g	enetic abn	ormalities iden	tified in those v	vith focal (par	tial) epilepsy				
17	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁴	6	2000	0.00 (0 to 0.01)	⊕000 VERY LOW	CRITICAL
Proportio	n of metabolic/g	enetic abn	ormalities iden	tified in those v	with Lennox-G	Sastaut syndro	ome			
1 ⁸	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁴	9	135	0.07 (0.03 to 0.12)	⊕000 VERY LOW	CRITICAL
Proportio	n of metabolic/g	enetic abn	ormalities iden	tified on 1.5-t						
29	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁴	4	399	0.01 (0 to 0.03)	⊕000 VERY LOW	CRITICAL
Proportio	n of metabolic/g	enetic abn	ormalities iden	tified on 3.0-t						

Quality as	Quality assessment					Number of patients Effect		Effect		
Number of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Number with a clinically relevant abnormality	Number assessed	Proportion (95% CI)	Quality	Importance
1 ¹⁰	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁴	6	2000	0.00 (0 to 0.01)	⊕000 VERY LOW	CRITICAL
Proportio	on of metabolic/g	enetic abn	ormalities in th	ose with a new	diagnosis					
3 ¹¹	Observational studies	Very serious ²	Serious ¹⁵	No serious indirectness	Very serious ⁴	23	1284	0.02 (0.01 to 0.04)	⊕000 VERY LOW	CRITICAL
Proportio	on of metabolic/g	enetic abn	ormalities iden	tified in those v	vith existing o	liagnosis and	treatmer	nt resistant		
1 ¹³	Observational studies	Very serious ²	No serious incosistency	No serious indirectness	Very serious ⁴	1	217	0.00 (0 to 0.03)	⊕000 VERY LOW	CRITICAL
Proportio	on of metabolic/g	enetic abn	ormalities iden	tified in those v	vithout learnii	ng disabilities	;			
18	Observational studies	Very serious ²	No serious incosistency	Serious ¹⁴	Very serious ⁴	9	135	0.07 (0.03 to 0.12)	⊕000 VERY LOW	CRITICAL
Proportio	Proportion of metabolic/genetic abnormalities identified in those who had a previous CT scan									

Quality a	ssessment					Number of p	atients	Effect		
Number of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Number with a clinically relevant abnormality	Number assessed	Proportion (95% CI)	Quality	Importance
1 ¹⁵	Observational studies	Very serious ²	No serious incosistency	No serious indirectness	Very serious ⁴	3	182	0.02 (0 to 0.05)	⊕000 VERY LOW	CRITICAL

- 1 Ali 2017, Asadi-Pooya 2012, Berg 2000, Bruno 2017, Coryell 2018, Craven 2012, Das 2013, Hsieh 2010, Petrou 2007
- 2 Very serious risk of bias in the evidence contributing to the outcomes as per CEBMA checklist
- 3 Very serious heterogeneity (*I*²>75%)
- 4 Number of events <150
- 5 Coryell 2018, Das 2013, Hsieh 2010, Petrou 2007
- 6 Berg 2000
- 7 Craven 2012
- 8 Asadi-Pooya 2012
- 9 Bruno 2017, Hsieh 2010
- 10 Craven 2012
- 11 Berg 200, Coryell 2018, Hsieh 2010
- 12 12>50% <75%
- 13 Bruno 2017
- 14 Population is indirect (3% of the participants did not have learning disabilities)
- 15 Hsieh 2010

Table 11: Clinical evidence profile for proportion identified with non-epilepsy related abnormalities

Quality a	Quality assessment					Number of patients		Effect		
Number of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Number with a clinically relevant abnormality	Number assessed	Proportion (95% CI)	Quality	Importance
Proportio	on identified with	non-epile	osy related abn	ormalities: ove	rall estimate					
201	Observational studies	Very serious ²	Very serious ³	No serious indirectness	No serious imprecision	700	6628	0.06 (0.04 to 0.09)	⊕000 VERY LOW	IMPORTANT
Proportio	on of non-epileps	sy related a	bnormalities id	entified in infa	nts (<3 years	old at seizure	onset)			
54	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Very serious ⁵	93	1421	0.08 (0.03 to 0.18)	⊕000 VERY LOW	IMPORTANT
Proportio	on of non-epileps	sy related a	bnormalities id	entified in child	dren (3 to 11 y	ears old at se	eizure ons	set)		
1 ⁶	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁵	15	388	0.04 (0.02 to 0.06)	⊕000 VERY LOW	IMPORTANT
Proportio	on of non-epileps	sy related a	bnormalities id	entified in adul	ts (25 to 65 ye	ears old sat s	eizure on	set)		
17	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁵	2	134	0.01 (0 to 0.05)	⊕000 VERY LOW	IMPORTANT

Quality as	uality assessment					Number of patients		Effect		
Number of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Number with a clinically relevant abnormality	Number assessed	Proportion (95% CI)	Quality	Importance
Proportio	n of non-epileps	y related a	bnormalities id	entified in thos	e with focal (partial) epilep	sy			
38	Observational studies	Very serious ²	Very serious ³	No serious indirectness	No serious imprecision	333	2183	0.07 (0.02 to 0.22)	⊕000 VERY LOW	IMPORTANT
Proportio	n of non-epileps	y related a	bnormalities id	entified in thos	e with genetic	c (idiopathic)	generalis	ed epilepsy		
5 ⁹	Observational studies	Very serious ²	Serious ¹⁰	No serious indirectness	Very serious ⁵	15	383	0.04 (0.02 to 0.10)	⊕000 VERY LOW	IMPORTANT
Proportio	n of non-epileps	y related a	bnormalities id	entified in thos	e with West s	yndrome				
111	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁵	0	2	0.00 (0 to 0.84)	⊕000 VERY LOW	IMPORTANT
Proportio	n of non-epileps	y related a	bnormalities id	entified in thos	e with Lenno	x-Gastaut syn	drome			
212	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	Very serious ⁵	1	137	0.01 (0 to 0.05)	⊕000 VERY LOW	IMPORTANT
Proportio	on of non-epileps	y related a	bnormalities id	entified on 1.5-	t					

Quality as	Quality assessment					Number of patients		Effect		
Number of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Number with a clinically relevant abnormality	Number assessed	Proportion (95% CI)	Quality	Importance
5 ¹³	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Very serious ⁵	78	688	0.10 (0.05 to 0.16)	⊕000 VERY LOW	IMPORTANT
Proportio	n of non-epileps	y related a	bnormalities id	entified on 3.0-	t					
114	Observational studies	Very serious ²	No serious inconsistency	No serious indirectness	No serious imprecision	326	2000	0.16 (0.15 to 0.18)	⊕000 VERY LOW	IMPORTANT
Proportio	n of non-epileps	y related a	bnormalities in	those with a n	ew diagnosis					
8 ¹⁵	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Serious ¹⁶	263	2733	0.06 (0.03 to 0.12)	⊕000 VERY LOW	IMPORTANT
Proportio	n of non-epileps	y related a	bnormalities id	entified in thos	e with existin	g diagnosis a	nd treatn	nent resistant		
217	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Very serious ⁵	28	311	0.01 (0.00 to 0.62)	⊕000 VERY LOW	IMPORTANT
Proportio	n of non-epileps	y related a	bnormalities id	entified in thos	e with existin	g diagnosis a	nd contro	olled		

Quality as	uality assessment					Number of p	atients	Effect		
Number of studies	Design	Risk of bias	Inconsistency	Indirectness	Imprecision	Number with a clinically relevant abnormality	Number assessed	Proportion (95% CI)	Quality	Importance
2 ¹⁸	Observational studies	Very serious ²	Serious ¹⁰	No serious indirectness	Very serious ⁵	8	202	0.05 (0.01 to 0.15)	⊕000 VERY LOW	IMPORTANT
Proportio	on of non-epileps	y related a	bnormalities id	entified in thos	e with learnin	g disabilities				
1 ¹⁹	Observational studies	Very serious ²	No serious inconsistency	Serious ²⁰	Very serious ⁵	1	135	0.01 (0 to 0.04)	⊕000 VERY LOW	IMPORTANT
Proportio	on of non-epileps	sy related a	bnormalities id	entified in thos	e without lear	rning disabilit	ies			
1 ²¹	Observational studies	Very serious ²	No serious inconsistency	Serious ²²	Very serious ⁵	4	32	0.12 (0.04 to 0.29)	⊕000 VERY LOW	IMPORTANT
Proportio	on of non-epileps	sy related a	bnormalities id	entified in thos	e who had a p	orevious CT s	can			
3 ²³	Observational studies	Very serious ²	Very serious ³	No serious indirectness	Very serious ⁵	40	383	0.07 (0.02 to 0.19)	⊕000 VERY LOW	IMPORTANT

¹ Ali 2017, Asadi-Pooya 2012, Aslan 2010, Bakhsh 2013, Benson 2009, Berg 2000, Betting 2006, Bruno 2017, Byars 2007, Coryell 2018, Craven 2012, Das 2013, Dirik 2018, Dura-Trave 2012, Ekici 2013, Hakami 2013, Hersdorffer 2008, Hsieh 2010, Rasool 2012, Wongladarom 2004

- 2 Very serious risk of bias in the evidence contributing to the outcomes as per CEBMA checklist
- 3 Very serious heterogeneity (I²>75%)
- 4 Hsieh 2010, Coryell 2018, Das 2013, Dirik 2018, Hesdorffer 2008
- 5 Number of events <150
- 6 Berg 2000
- 7 Betting 2006
- 8 Craven 2012, Rasool 2012, Wongladarom 2004
- 9 Aslan 2010, Bakhsh 2013, Betting 2006, Rasool 2012, Wongladarom 2004
- 10 Serious heterogeneity (I² >50% but <75%)
- 11 Wongladarom 2004
- 12 Asadi-Pooya 2012, Wongladarom 2004
- 13 Asaln 2010, Bruno 2017, Hsieh 2010, Rasool 2012, Wongladarom 2004
- 14 Craven 2012.
- 15 Benson 2019, Berg 2000, Byars 2007, Coryell 2018, Dirik 2018, Rasool 2012, Hakami 2013, Hsieh 2010
- 16 Number of events >150 but <300
- 17 Bruno 2017, Ekici 2013
- 18 Aslan 2010, Ekici 2013
- 19 Asadi-Pooya 2012
- 20 Population is indirect in 1 study (3% of participants did not have learning disabilities)
- 21 Aslan 2010
- 22 Population is indirect in 1 study (3% of participants did have learning disabilities)
- 23 Bakshsh 2013, Hsieh 2010, Rasool 2012

Appendix G – Economic evidence study selection

Economic evidence study selection for review question: What is the effectiveness of genetic testing in determining the aetiology of epilepsy?

A global search of economic evidence was undertaken for all review questions in this guideline. See Supplement 2 for further information

Appendix H – Economic evidence tables

Economic evidence tables for review question: What is the effectiveness of genetic testing in determining the aetiology of epilepsy?

No evidence was identified which was applicable to this review question

Appendix I – Economic evidence profiles

Economic evidence profiles for review question: What is the effectiveness of genetic testing in determining the aetiology of epilepsy?

No evidence was identified which was applicable to this review question

Appendix J - Economic analysis

Economic evidence analysis for review question: What is the yield of relevant abnormalities detected by MRI in people with epilepsy?

No economic analysis was conducted for this review question.

Appendix K – Excluded studies

Excluded clinical and economic studies for review question: What is the yield of relevant abnormalities detected by MRI in people with epilepsy?

Clinical studies

Table 12: Excluded studies and reasons for their exclusion

able 12: Excluded studies and reasons for t Excluded studies - Yield of MRI	
Study	Reason for Exclusion
Aamir, I., Arooj, S., Mansoor, M., Niazi, T., Neuroimaging in epilepsy: Magnetic resonance imaging (MRI) evaluation in refractory complex partial seizures, Pakistan Journal of Medical and Health Sciences, 8, 1105-1108, 2014	No relevant study design; case series
Adachi, Y., Yagishita, A., Arai, N., White matter abnormalities in the anterior temporal lobe suggest the side of the seizure foci in temporal lobe epilepsy, Neuroradiology, 48, 460-464, 2006	Yield of MRI abnormalities was not reported
Adams, M. E., Aylett, S. E., Squier, W., Chong, W., A Spectrum of unusual neuroimaging findings in patients with suspected Sturge-Weber syndrome, American Journal of Neuroradiology, 30, 276-281, 2009	Incorrect population
Agarwal, A., Raghav, S., Husain, M., Kumar, R., Gupta, R. K., Epilepsy with focal cerebral calcification: Role of magnetization transfer MR imaging, Neurology India, 52, 197-199, 2004	No relevant study design; case control study
Alhusaini, S., Doherty, C. P., Scanlon, C., Ronan, L., Maguire, S., Borgulya, G., Brennan, P., Delanty, N., Fitzsimons, M., Cavalleri, G. L., A cross-sectional MRI study of brain regional atrophy and clinical characteristics of temporal lobe epilepsy with hippocampal sclerosis, Epilepsy Research, 99, 156-166, 2012	No relevant outcomes were reported; the study described MRI-based volumetric analysis
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Andres, M., Andre, V. M., Nguyen, S., Salamon, N., Cepeda, C., Levine, M. S., Leite, J. P., Neder, L., Vinters, H. V., Mathern, G. W., Human cortical dysplasia and epilepsy: An ontogenetic hypothesis based on volumetric MRI and NeuN neuronal density and size	Incorrect diagnostic test

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Aprahamian, N., Harper, M. B., Prabhu, S. P., Monuteaux, M. C., Sadiq, Z., Torres, A., Kimia, A. A., Pediatric first time non-febrile seizure with focal manifestations: Is emergent imaging indicated?, Seizure, 23, 740-745, 2014	CT and MRI were performed, but results have not been reported separately
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Arya, R., Mangano, F. T., Horn, P. S., Kaul, S. K., Roth, C., Leach, J. L., Turner, M., Holland, K. D., Greiner, H. M., Long-term seizure outcomes after pediatric temporal lobectomy: Does brain MRI lesion matter?, Journal of Neurosurgery: Pediatrics, 24, 200-208, 2019	Yield of MRI abnormalities was not reported
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Basiri, R., Shariatzadeh, A., Wiebe, S., Aghakhani, Y., Focal epilepsy without interictal spikes on scalp EEG: A common finding of uncertain significance, Epilepsy Research, 150, 1-6, 2019	Yield of MRI abnormalities was not reported
Bayram, E., Topcu, Y., Yis, U., Cakmaci, H., Kurul, S. H., Comparison of cranial magnetic resonance imaging findings and clinical features in patients with corpus callosum abnormalities, Neuropediatrics, 45, 30-35, 2014	Not all patients presented with epilepsy and the results could not be extracted for the target population
Bekelis, K., Desai, A., Kotlyar, A., Thadani, V., Jobst, B. C., Bujarski, K., Darcey, T. M., Roberts, D. W., Occipitotemporal hippocampal depth electrodes in intracranial epilepsy monitoring: Safety and utility; Clinical article, Journal of Neurosurgery, 118, 345-352, 2013	Proportion of specific abnormalities was not reported
Berger, J., Plotkin, M., Demin, K., Holtkamp, M., Bengner, T., The relationship between structural MRI, FDG-PET, and memory in temporal lobe epilepsy: Preliminary results, Epilepsy and Behavior, 80, 61-67, 2018	No relevant outcomes were reported
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Classification and prognostics, Annals of	
Neurology, 77, 436-446, 2015	
Bersani, G., Iannitelli, A., Quartini, A., Di Biasi,	Not an investigation of a standardised MRI
C., Gualdi, G., Pancheri, P., Patients with epilepsy associated with schizophrenia: A	programme
descriptive study of patients investigated with	
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standard electroencephalography (EEG), Italian	
Journal of Psychopathology, 14, 10-15, 2008	
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development and incomplete hippocampal inversion with medically intractable seizures in	
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Prevalence and distribution of MRI abnormalities	abnormality, only its location
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H., Mateen, F. J., Neurocysticercosis in Bhutan:	
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Calado, S., Jordao, C., Vale, J., The	Study design not relevant, case series
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Citterio, A., Sberna, M., Tassi, L., Mai, R.,	·
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Russo, G., Colombo, N., SUrface-PRojected	
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A Novel Tool for Advanced Imaging of Epilepsy,	
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I., Helmstaedter, C., von Oertzen, J., Urbach, H.,	
Schramm, J., Lesional mesial temporal lobe	
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Clusmann, H., Schramm, J., Kral, T.,	No relevant outcomes were reported
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outcome after different types of resection for	
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Dakaj, N., Kruja, J., Jashari, F., Boshnjaku, D.,	Study does not report the yield of MRI
Shatri, N., Zeqiraj, K., Accuracy of conventional	abnormalities
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Bartolini, E., Biagi, L., Cossu, M., Pelliccia, V.,	
Symms, M. R., Guerrini, R., 7T MRI in focal	
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drug-resistant epilepsy due to focal cortical	
dysplasia type 2: additional value of	
electroclinical data and coregistration with MRI,	
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Molecular Imaging, 45, 1449-1460, 2018	
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Hamartomas and epilepsy: Clinical and imaging	
characteristics, Seizure, 12, 307-311, 2003	
Ding, Y. S., Chen, B. B., Glielmi, C., Friedman,	Not relevant study design; case series
K., Devinsky, O., A pilot study in epilepsy	
patients using simultaneous PET/MR, American	
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Imaging, 4, 459-470, 2014	Viold of MPI abnormalities was not reported
Doescher, J. S., deGrauw, T. J., Musick, B. S., Dunn, D. W., Kalnin, A. J., Egelhoff, J. C.,	Yield of MRI abnormalities was not reported
Bryars, A. W., Mathews, V. P., Austin, J. K.,	
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electroencephalographic (EEG) findings in a	
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209-213, 2016	

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Eeg-Olofsson, O., Lundberg, S., Raininko, R.,	Conference abstract
MRI in rolandic epilepsy, Epileptic Disorders, 2 Suppl 1, S51-3, 2000	Conference abstract
El Ameen, N. F., Amin, M. F., kotb, A., MRI of	Patients did not present with epilepsy
the brain in postpartum convulsions; pose	1 1 7
diagnostic dilemmas, Egyptian Journal of	
Radiology and Nuclear Medicine, 48, 999-1004,	
2017	
Farrow, T. F. D., Dickson, J. M., Grunewald, R.	No relevant study design; case series
A., A Six-Year Follow-Up MRI Study of	·
Complicated Early Childhood Convulsion,	
Pediatric Neurology, 35, 257-260, 2006	
Fredriksen, J. R., Carr, C. M., Koeller, K. K.,	Irrelevant study design; case series
Verdoorn, J. T., Gadoth, A., Pittock, S. J.,	
Kotsenas, A. L., MRI findings in glutamic acid	
decarboxylase associated autoimmune epilepsy,	
Neuroradiology, 60, 239-245, 2018	
Gaily, E., Anttonen, A. K., Valanne, L.,	No relevant study design; case series
Liukkonen, E., Traskelin, A. L., Polvi, A., Lommi,	
M., Muona, M., Eriksson, K., Lehesjoki, A. E.,	
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modifiers, imaging abnormalities, and ictal	
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Gilliam, F., Faught, E., Martin, R., Bowling, S., Bilir, E., Thomas, J., Morawetz, R., Kuzniecky,	No relevant outcomes were reported
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temporal lobe epilepsy: An intent-to-treat	
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Glass, H. C., Bonifacio, S. L., Sullivan, J.,	Population were newborn babies
Rogers, E., Ferriero, D. M., Goldstein, R.,	r opalation were newsom suspec
Barkovich, J. A., Magnetic resonance imaging	
and ultrasound injury in preterm infants with	
seizures, Journal of Child Neurology, 24, 1105-	
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Goyal, M., Bangert, B. A., Lewin, J. S., Cohen,	No relevant study design; case series
M. L., Robinson, S., High-resolution MRI	
enhances identification of lesions amenable to	
surgical therapy in children with intractable	
epilepsy, Epilepsia, 45, 954-959, 2004	
Grillo, E., Postictal MRI abnormalities and	No relevant outcomes were reported
seizure-induced brain injury: Notions to be	
challenged, Epilepsy and Behavior, 44, 195-199, 2015	
	No relevant outcomes were reported
Grunewald, R. A., Farrow, T., Vaughan, P., Rittey, C. D. C., Mundy, J., A magnetic	No relevant outcomes were reported
resonance study of complicated early childhood	
convulsion, Journal of Neurology Neurosurgery	
and Psychiatry, 71, 638-642, 2001	
Gunawan, P. I., Saharso, D., Purnama Sari, D.,	No relevant outcomes were reported
Correlation of serum S100B levels with brain	The relevant editedines were reported
magnetic resonance imaging abnormalities in	
children with status epilepticus, Korean Journal	
of Pediatrics, 62, 281-285, 2019	
Gupta, S. N., Belay, B., Intracranial incidental	The study does not specify whether all included
findings on brain MR images in a pediatric	patients had epilepsy
neurology practice: A retrospective study,	
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Sci, 264, 34-37, 2008	

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Halac, G., Delil, S., Zafer, D., Isler, C., Uzan, M., Comunoglu, N., Oz, B., Yeni, S. N., Vatankulu, B., Halac, M., Ozkara, C., Compatibility of MRI and FDG-PET findings with histopathological results in patients with focal cortical dysplasia, Seizure, 45, 80-86, 2017	No relevant outcomes were reported
Hallbook, T., Ruggieri, P., Adina, C., Lachhwani, D. K., Gupta, A., Kotagal, P., Bingaman, W. E., Wyllie, E., Contralateral MRI abnormalities in candidates for hemispherectomy for refractory epilepsy, Epilepsia, 51, 556-563, 2010	No relevant outcomes were reported
Heers, M., Rampp, S., Stefan, H., Urbach, H., Elger, C. E., von Lehe, M., Wellmer, J., MEGbased identification of the epileptogenic zone in occult peri-insular epilepsy, Seizure, 21, 128-33, 2012	No relevant outcomes were reported
Ho, K., Lawn, N., Bynevelt, M., Lee, J., Dunne, J., Neuroimaging of first-ever seizure Contribution of MRI if CT is normal, Neurology: Clinical Practice, 3, 398-403, 2013	CT and MRI were performed, but results have not been reported separately
Izuora, G. I., Ayadi, K. M., Okoroma, E., Neuroimaging findings in children with infantile spasms, Neurosciences, 9, 30-33, 2004	No relevant study design; case series
Jahodova, A., Krsek, P., Kyncl, M., Jezdik, P., Kudr, M., Komarek, V., Jayakar, P., Miller, I., Resnick, T., Duchowny, M., Distinctive MRI features of the epileptogenic zone in children with tuberous sclerosis, European Journal of Radiology, 83, 703-709, 2014	No relevant outcomes were reported
Jansen, J. F. A., Vlooswijk, M. C. G., Majoie, H. M., De Krom, M. C. T. F. M., Aldenkamp, A. P., Hofman, P. A. M., Backes, W. H., White matter lesions in patients with localization-related epilepsy, Investigative Radiology, 43, 552-558, 2008	No relevant outcomes were reported
Kalnin, A. J., Fastenau, P. S., deGrauw, T. J., Musick, B. S., Perkins, S. M., Johnson, C. S., Mathews, V. P., Egelhoff, J. C., Dunn, D. W., Austin, J. K., Magnetic Resonance Imaging Findings in Children With a First Recognized Seizure, Pediatric Neurology, 39, 404-414, 2008	Unable to read the contents of the Appendix where the results were reported as these were distorted. Author was contacted, but no response received
Kasasbeh, A., Hwang, E. C., Steger-May, K., Bandt, S. K., Oberhelman, A., Limbrick, D., Miller-Thomas, M. M., Shimony, J. S., Smyth, M. D., Association of magnetic resonance imaging identification of mesial temporal sclerosis with pathological diagnosis and surgical outcomes in children following epilepsy surgery: Clinical article, Journal of Neurosurgery: Pediatrics, 9, 552-561, 2012	Irrelevant study design; case series
Katramados, A. M., Burdette, D., Patel, S. C., Schultz, L. R., Gaddam, S., Mitsias, P. D., Periictal diffusion abnormalities of the thalamus in partial status epilepticus, Epilepsia, 50, 265-75, 2009	Irrelevant study design; case series
Kim, D. W., Lee, S. K., Yun, C. H., Kim, K. K., Lee, D. S., Chung, C. K., Chang, K. H., Parietal lobe epilepsy: The semiology, yield of diagnostic workup, and surgical outcome, Epilepsia, 45, 641-649, 2004	Irrelevant study design; case series

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Lascano, A. M., Perneger, T., Vulliemoz, S., Spinelli, L., Garibotto, V., Korff, C. M., Vargas, M. I., Michel, C. M., Seeck, M., Yield of MRI, high-density electric source imaging (HD-ESI), SPECT and PET in epilepsy surgery candidates, Clinical Neurophysiology, 127, 150-155, 2016	Yield of MRI was not reported
Lefkopoulos, A., Tzinas, A., Papadopoulou, E., Haritanti, A., Karanikolas, D., Tsifountoudis, I., Dimitriadis, A. S., MRI assessment of hippocampal sclerosis, Rivista di Neuroradiologia, 18, 357-363, 2005	Irrelevant study design; case series
Liu, R. S. N., Lemieux, L., Bell, G. S., Bartlett, P. A., Sander, J. W. A. S., Sisodiya, S. M., Shorvon, S. D., Duncan, J. S., A longitudinal quantitative MRI study of community-based patients with chronic epilepsy and newly diagnosed seizures: Methodology and preliminary findings, Neurolmage, 14, 231-243, 2001	Conference abstract
Liu, R. S. N., Lemieux, L., Bell, G. S., Sisodiya, S. M., Bartlett, P. A., Shorvon, S. D., Sander, J. W. A. S., Duncan, J. S., Cerebral damage in epilepsy: A population-based longitudinal quantitative MRI study, Epilepsia, 46, 1482-1494, 2005	No relevant outcomes were reported
Lizcano, A., Carrico, L., Barbosa, P., Carvalho, M. I., Yasuda, C., Montenegro, M. A., Guerreiro, M., Guerreiro, C., Cendes, F., EEG and magnetic resonance imaging abnormalities in patients with acute limbic encephalitis, Journal of Epilepsy and Clinical Neurophysiology, 17, 133-139, 2011	Not relevant study design; case series
Lyons, T. W., Johnson, K. B., Michelson, K. A., Nigrovic, L. E., Loddenkemper, T., Prabhu, S. P., Kimia, A. A., Yield of emergent neuroimaging in children with new-onset seizure and status epilepticus, Seizure, 35, 4-10, 2016	Not relevant study design; case series
Malik, M. A., Tarar, M. A., Hamid, H., Ur Rehhman, M., Qureshi, A., Ossaid, M., Sultan, T., Ahmad, N., Ali, Q., Malik, S., Diagnostic importance of interictal electroencephalogram and neuroimaging of brain in new-onset idiopathic generalized epilepsy of childhood (IGEC), Pakistan Paediatric Journal, 34, 15-22, 2010	Unavailable. Last checked 29/03/21
Marsh, L., Sullivan, E. V., Morrell, M., Lim, K. O., Pfefferbaum, A., Structural brain abnormalities in patients with schizophrenia, epilepsy, and epilepsy with chronic interictal psychosis, Psychiatry Research, 108, 1-15, 2001	Mixed population of people with epilepsy and schizophrenia. Results were not reported separately
Matsuura, K., Maeda, M., Okamoto, K., Araki, T., Miura, Y., Hamada, K., Kanamaru, K., Tomimoto, H., Usefulness of arterial spinlabeling images in periictal state diagnosis of epilepsy, Journal of the Neurological Sciences, 359, 424-429, 2015	No relevant outcomes were reported
McGill, M. L., Devinsky, O., Wang, X., Quinn, B. T., Pardoe, H., Carlson, C., Butler, T., Kuzniecky, R., Thesen, T., Functional neuroimaging abnormalities in idiopathic	No relevant outcomes were reported

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generalized epilepsy, NeuroImage: Clinical, 6,	
455-462, 2014	
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P., Neuroimaging in Epilepsy, Current Neurology	Ivaliative review
and Neuroscience Reports, 17 (4) (no	
pagination), 2017	
Milligan, T. A., Zamani, A., Bromfield, E.,	No relevant study design; case series
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2009	
Mitsueda-Ono, T., Ikeda, A., Sawamoto, N.,	No relevant study design; case series
Aso, T., Hanakawa, T., Kinoshita, M.,	
Matsumoto, R., Mikuni, N., Amano, S., Fukuyama, H., Takahashi, R., Internal structural	
changes in the hippocampus observed on 3-	
tesla MRI in patients with mesial temporal lobe	
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Morimoto, E., Kanagaki, M., Okada, T.,	Participants did not have epilepsy
Yamamoto, A., Mori, N., Matsumoto, R., Ikeda,	
A., Mikuni, N., Kunieda, T., Paul, D., Miyamoto,	
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Wheless, J. W., Carmant, L., Bebin, M., Conry, J. A., Chiron, C., Elterman, R. D., Frost, M.,		
J. A., Chiron, C., Elterman, R. D., Frost, M., reported		Yield of specific MRI abnormalities was not
	Paolicchi, J. M., Donald Shields, W., Thiele, E.	

Excluded studies - Yield of MRI	
A., Zupanc, M. L., Collins, S. D., Magnetic resonance imaging abnormalities associated with vigabatrin in patients with epilepsy, Epilepsia, 50, 195-205, 2009	
Whiting, P., Gupta, R., Burch, J., Mota, R. E., Wright, K., Marson, A., Weishmann, U., Haycox, A., Kleijnen, J., Forbes, C., A systematic review of the effectiveness and cost-effectiveness of neuroimaging assessments used to visualise the seizure focus in people with refractory epilepsy being considered for surgery, Health technology assessment (Winchester, England), 10, 1-250, iii-iv, 2006	No relevant outcomes were reported
Widjaja, E., Nilsson, D., Blaser, S., Raybaud, C., White matter abnormalities in children with idiopathic developmental delay, Acta Radiologica, 49, 589-95, 2008	Not all patients presented with epilepsy and the results could not be extracted for the target population
Widjaja, E., Otsubo, H., Raybaud, C., Ochi, A., Chan, D., Rutka, J. T., Snead, lii O. C., Halliday, W., Sakuta, R., Galicia, E., Shelef, I., Chuang, S. H., Characteristics of MEG and MRI between Taylor's focal cortical dysplasia (type II) and other cortical dysplasia: Surgical outcome after complete resection of MEG spike source and MR lesion in pediatric cortical dysplasia, Epilepsy Research, 82, 147-155, 2008	Study does not report the yield of MRI abnormalities, only its location
Wychowski, T., Hussain, A., Tivarus, M. E., Birbeck, G. L., Berg, M. J., Potchen, M., Qualitative analysis of double inversion recovery MRI in drug-resistant epilepsy, Epilepsy Research, 127, 195-199, 2016	No relevant outcomes were reported
Xiang, T., Li, G., Liang, Y., Zhou, J., A wide spectrum of variably periictal MRI abnormalities induced by a single or a cluster of seizures, Journal of the Neurological Sciences, 343, 167-172, 2014	No relevant outcomes were reported

Economic studies

A global search of economic evidence was undertaken for all review questions in this guideline. See Supplement 2 for further information.

Appendix L – Research recommendations

Research recommendations for review question:

No research recommendations were made for this review question.

Appendix M - Clinically relevant abnormalities

Clinically relevant abnormalities have been categorised as follows:

- Tumour
 - o Brain metastases
 - o Primary brain tumours, including meningiomas
- Vascular
 - o Arterio-venous malformation (AVM)/vascular malformation/abnormality
 - o Haemorrhage
 - Infarct/ Infarction
 - o PRES (posterior reversible encephalopathy syndrome)
 - Vasculitis
 - Venous sinus thrombosis
- Scarring
 - o Encephalomalacia/cystic encephalomalacia
 - Gliosis
 - o Hippocampal sclerosis/ Mesial temporal sclerosis
 - o Ulegyria
- Congenital/developmental
 - Dysmyelination
 - o Hydrocephalus
 - o Malformations of cortical development
 - o Phakomatoses
- Inflammatory/infective/immune
 - o Autoimmune encephalitis/limbic encephalitis
 - o Demyelination
 - o Infections
 - o Oedema/edema
- Metabolic /Genetic
 - o Congenital disorders of glycosylation/Carbohydrate deficient glycoprotein disorders
 - Disorders of amino acid metabolism
 - Glucose transporter deficiency
 - Leucodystrophy (including very long chain fatty acid disorders)
 - Lysosomal enzyme disorders
 - Mitochondrial Disorders
 - Molybdenum cofactor deficiency
 - Organic acidurias
 - Sulphite oxidase deficiency