Ranking	Uncertainty	Additional notes	Source url:	Suggested Outcomes	Outcomes to be measured	Reference to up-to date systematic review	Reference to out of date SR	Reference to an up-to date, relevant and reliable SR which does not refer to uncertainty but could be extended	Protocol for a systematic review	Ongoing studies	Comparison	Health Topic	Health Topic	Uncertainty from Patient/Carer/Clinician/R esearch Recommendation or multiple sources	Age of patients/population
1	Does treatment (with neurosurgery or stereotactic radiosurgery) or no treatment improve outcome for people diagnosed with brain or spine cavernoma?	three others presented at the PSP workshop (11, 19 and 20, below). Survey carried out by	0.interdns.co.uk/our- , projects-and- campaigns/priority-	Prognosis: change in symptoms, or change in management of symptoms: adverse effects or complications; patient satisfaction; health related quality of life; service related issues; and health related cost	Change of symptoms; acceptability to patients	Wang X, Mei G, Liu X, Dai J, Pan L, Wang E. The role of stereotactic radiosurgery in cavernous sinus hemangiomas: a systematic review and meta-analysis. J Neuro-Oncol. 2012;107(2):239-45. doi: 10.1007/s11060-011-0753-8. Epub 2011 Nov 16. Lu XY, Sun H, Xu JG, Li QY. Stereotactic radiosurgery of brainstem cavernous malformations: a systematic review and meta-analysis. J Neurosurg. 2014;120(4):982-7. doi: 10.3171/2013.12.JNS13990. Epub 2014 Feb 7. Badhiwala J.H. Farrokhya F. Alhazzani W. Yarascavitch B. Ed Aref M. Algird A, et al. Surgical outcomes and natural history of intramedullary spinal cord cavernous malformations: a single-center series and meta- analysis of individual patient data. Journal of Neurosurgery: Spine. 2014;21 (4):662-76. doi: 10.3171/2014.6.SPINE13949. Epub 2014 Jul 25. Poorthuis M. Samarasekera N. Kontoh K. Stuart I. Cope B. Kitchen N. Al-Shahi Salman R. Comparative studies of the diagnosis and treatment of cerebral cavernous malformations in adults: Systematic review Acta Neurochirurg. 2013;155 (4):643-9. doi: 10.1007/S00710-13-1621-4. Epub 2013 Jan 31. Poorthuis MH, Klijn CJ, Algra A, Rinkel GJ, Al-Shahi Salman R. Treatment of cerebral cavernous malformations: a systematic review and meta-regression analysis. J Neurol Neurosurg Psychiatr. 2014; 851(2):1319-23. doi: 10.1136/jnnp- 2013-307349. Epub 2014 Mar 25.		n/a	n/a	n/a	Surgery: Service delivery	Neurological conditions:Brain and spinal cavernomas	Neurological conditions Surgery, anaesthesia, perioperative and critical care	120 patients. 28 clinicians.	Any age
2	How do brain or spine cavernomas start and develop?	Survey carried out by the Cavernom Alliance UK for this PSP.	http://cavernoma1.users4 0.interdns.co.uk/our- projects-and- campaigns/priority- setting-partnership-psp- project/	Aetiology	Aetiology	n/a	n/a	n/a	n/a	n/a	Aetiology	Neurological conditions:Brain and spinal cavernomas	Neurological conditions	90 patients. 20 carers.	Any age
3	What is the risk of brain or spine cavernomas bleeding for the first and subsequent times?	Survey carried out by the Cavernom Alliance UK for this PSP.	http://cavernoma1.users4 0.interdns.co.uk/our- projects-and- campaigns/priority- setting-partnership-psp- project/	Prognosis	Adverse effects or complications	n/a	n/a	n/a	n/a	n/a	Prognosis	Neurological conditions:Brain and spinal cavernomas	Neurological conditions	113 patients.	Any age
4	Could drugs targeted at cavernomas improve outcome for people with brain or spine cavernomas compared to no drug treatment?	Survey carried out by the Cavernom Alliance UK for this PSP.	0.interdns.co.uk/our- projects-and- campaigns/priority-	Prognosis: change in symptoms, or change in management of symptoms; adverse effects or complications; patient satisfaction; health related quality of life; service related issues; and health related rost	Change of symptoms	n/a	n/a	n/a	n/a	n/a	Prognosis	Neurological conditions:Brain and spinal cavernomas	Neurological conditions	45 patients.	Any age
5	What mechanisms trigger bleeding or epileptic seizures in people with brain or spine cavemornas?	Survey carried out by the Cavernom Alliance UK for this PSP.	http://cavernoma1.users4 0.interdns.co.uk/our- projects-and- campaigns/priority- setting-partnership-psp- project/	Aetiology	Adverse effects or complications; health related quality of life	n/a	n/a	n/a	n/a	n/a	Aetiology	Neurological conditions:Brain and spinal cavernomas; Neurological conditions:Epilepsy	Neurological conditions Epilepsy	45 patients. 5 carers.	Any age
6	Are any features of brain or spine cavernoma on imaging associated with a higher or lower risk of bleeding?	Survey carried out by the Cavernom Alliance UK for this PSP.	0.interdns.co.uk/our- projects-and- campaigns/priority-	Diagnostic: prognosis; change in symptoms, or change in management of symptoms; adverse effects or complications; patient satisfaction; health related quality of life; service related issues; and health related cost	Change of symptoms; Adverse effects or complications; acceptability to patients or carers	n/a	n/a	n/a	n/a	n/a	Aetiology; Prognosis	Neurological conditions:Brain and spinal cavernomas	Neurological conditions Diagnostics	12 patients.	Any age
7	Does the use of anticoagulant drugs increase the risk of bleeding from brain or spine cavernoma?	Survey carried out by the Cavernom Alliance UK for this PSP.	0.interdns.co.uk/our- projects-and- campaigns/priority-	Prognosis: change in symptoms, or change in management of symptoms; adverse effects or complications; patient satisfaction; health related quality of life; service related issues; and health related cost	Change of symptoms; acceptability to patients or carers	n/a	n/a	n/a	n/a	n/a	Aetiology; Prognosis	Neurological conditions:Brain and spinal cavernomas	Neurological conditions Drug	45 patients. 7 clinicians.	Any age
8	Does regular monitoring of brain or spine cavernoma improve outcome compared to no monitoring?	Survey carried out by the Cavernom Alliance UK for this PSP.	0.interdns.co.uk/our- projects-and- campaigns/priority-	Diagnostic: prognosis; change in symptoms, or change in management of symptoms; adverse effects or complications; patient satisfaction; health related quality of life; service related issues; and health related cost	Change of symptoms	n/a	n/a	Na	n/a	n/a	Aetiology; Diagnostic; Prognosis; Service delivery	Neurological conditions:Brain and spinal cavernomas	Neurological conditions diagnostics	40 patients. 13 clinicians.	Any age
9	What features of brain cavernoma lead to the development of epilepsy, or influence the severity of existing epilepsy?	Existing guideline.	om/doi/10.1111/epi.1240 2/abstract;jsessionid=EE 2DBD13C870DCFC3FA D26F1BEDD44BD.f01t0 1	Aetiology: prognosis: change in symptoms, or change in management of symptoms; adverse effects or complications; patient satisfaction; health related quality of life; service related issues; and health related cost	Change of symptoms; health related quality of life	Epilepsia. 2013;54(12): 2025-35. DOI:	Shorvon S. Kendall BE. Stevens JM. Supratentorial cavernous haemangiomas and epilepsy: a review of the literature and case series. Neurosurg Psychiatr. 1999;	seizure(s) associated with cerebral cavernous malformations or arteriovenous malformations. Epilepsia. 2012;53 (SUPPL. 4):34-42. doi	;	n/a	Aetiology; Prognosis	Neurological conditions:Brain and spinal cavernomas; Neurological conditions:Epilepsy	Neurological conditions epilepsy	Research recommendation	Any age
10	Do any specific activities undertaken by people with brain or spine cavernomas provoke bleeds or other symptoms?	Survey carried out by the Cavernom Alliance UK for this PSP.	0.interdns.co.uk/our- projects-and- campaigns/priority-	Prognosis: change in symptoms, or change in management of symptoms; adverse effects or complications; patient satisfaction; health related quality of life; service related issues; and health related cost	Adverse effects or complications; acceptability to patients or carers; health related quality of life		n/a	n/a	n/a	n/a	Prognosis	Neurological conditions:Brain and spinal cavernomas	Neurological conditions Exercise	200 patients. 74 carers.	Any age

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Image: Problem in the state of the state	What is the impact of brain or spine cavernomas on life expectancy?			Prognosis		n/a	n/a	n/a	n/a n/a	Prognosis		Neurological conditions	15 patients. 3 carers.	Any age
		PSP.			related quality of me									
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Image: A large problem in the second problem in				life; service related issues; and										
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Image: Note of the second s	Why do only around half of people with a cavernoma gene mutation develop symptoms			Prognosis		n/a	n/a	n/a	n/a n/a	Aetiology; Prognosis	Neurological	Neurological conditions	60 patients. 28 carers.	Any age
And the set of			0.interdns.co.uk/our-		hospital and or needing health or social						conditions:Brain and	-		
Address and sectors and s		r'or.			care services, nearm related quality of life						spinai cavernomas			
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Image: Control (Control (Contro(Control (Control (Control (Control (Control (Control (Control (Co	What causes brain or spine cavernomas arising following radiotherapy for brain tumour			Aetiology: prognosis	Adverse effects or complications;	n/a	n/a	n/a	n/a n/a	Aetiology; Prognosis	Neurological		4 patients.	Any age
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Image: Section of the sectin of the section of the section	What causes single brain or spine cavernomas?	Survey carried out by the	http://cavernoma1.users4	Aetiology	Aetiology	n/a	n/a	n/a	n/a n/a	Aetiology	Neurological	Neurological conditions	3 patients.	Any age
Image: Problem Proble		Cavernom Alliance UK for this	0.interdns.co.uk/our-								conditions:Brain and	0		, ,
		PSP.									spinal cavernomas			
Maximum protection pr														
Maximum protection pr	Why do some patients develop multiple cavernomas even though they do not have any	of Survey carried out by the	project/ http://cavernoma1.users4	Aetiology	Aetiology	n/a	n/a	n/a	n/a n/a	Aetiology; Prognosis	Neurological	Neurological conditions	4 patients. 1 clinician.	Any age
Image: A problem of the state of the sta			0.interdns.co.uk/our-							0,. 0	conditions:Brain and			, ,
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Image: Note the set of the set o	What are the non-genetic causes of brain or spine cavernomas?	Cavernom Alliance UK for this	project/ http://cavernoma1.users4 0.interdns.co.uk/our-	Aetiology	Aetiology	n/a	n/a	n/a	ı/a n/a	Aetiology	conditions:Brain and	Neurological conditions	20 patients. 7 carers.	Any age
Methodskale Statuskale Status	What are the non-genetic causes of brain or spine cavernomas?	Cavernom Alliance UK for this	project/ http://cavernoma1.users4 0.interdns.co.uk/our- projects-and-	Aetiology	Aetiology	n/a	n/a	n/a	ı/a n/a	Aetiology	conditions:Brain and	Neurological conditions	20 patients. 7 carers.	Any age
Parameters Description control Description control Parameters	What are the non-genetic causes of brain or spine cavernomas?	Cavernom Alliance UK for this	project/ http://cavernoma1.users4 0.interdns.co.uk/our- projects-and- campaigns/priority- setting-partnership-psp-	Aetiology	Aetiology	n/a	n/a	n/a	v/a n/a	Aetiology	conditions:Brain and	Neurological conditions	20 patients. 7 carers.	Any age
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Epub 2014 Mar 25.	n/a	n/a	va n/a va n/a va n/a va n/a va n/a va n/a	Psychological therapy Aetiology; Prognosis Service delivery Aetiology; Prognosis Service delivery Prognosis	conditions:Brain and spinal cavernomas Neurological conditions:Brain and spinal cavernomas; Mental Health Neurological conditions:Brain and spinal cavernomas Neurological conditions:Brain and spinal cavernomas Neurological conditions:Brain and spinal cavernomas Neurological conditions:Brain and spinal cavernomas Neurological conditions:Brain and spinal cavernomas	Neurological conditions; Neurological conditions; diagnostics Neurological conditions; Service delivery Neurological conditions; Service delivery Neurological conditions; Social care Neurological conditions; Social care Neurological conditions; Surgery; Radiotherapy Neurological conditions;	2 clinicians. 20 patients. 3 clinicians. 5 patients. 3 carers. 25 patients. 15 carers. 25 patients. 15 carers. Multiple sources.	Any age
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Epub 2014 Mar 25.	n/a n/a n/a n/a n/a n/a n/a n/a n/a	n/a	va n/a va n/a va n/a va n/a va n/a va n/a	Psychological therapy Aetiology; Prognosis Service delivery Aetiology; Prognosis Service delivery Prognosis	conditions:Brain and spinal cavernomas Neurological conditions:Brain and spinal cavernomas; Mental Health Neurological conditions:Brain and spinal cavernomas Neurological conditions:Brain and spinal cavernomas Neurological conditions:Brain and spinal cavernomas Neurological conditions:Brain and spinal cavernomas Neurological conditions:Brain and spinal cavernomas	Neurological conditions; Neurological conditions; diagnostics Neurological conditions; Service delivery Neurological conditions; Service delivery Neurological conditions; Social care Neurological conditions; Social care Neurological conditions; Surgery; Radiotherapy Neurological conditions;	2 clinicians. 20 patients. 3 clinicians. 5 patients. 3 carers. 25 patients. 15 carers. 25 patients. 15 carers. Multiple sources.	Any age

26	Is it possible to identify signs and symptoms that are unique to brain or spine	Survey carried out by the		Adverse effects or complications;	n/a	n/a	n/a	n/a	n/a	Aetiology; Prognosis	Neurological	Neurological conditions;	8 patients. 8 clinicians.	Any age
	cavernomas?	Cavernom Alliance UK for this									conditions:Brain and	diagnostics		
		PSP.	projects-and-								spinal cavernomas			
			campaigns/priority-											
			setting-partnership-psp- project/											
27	What is the optimum radiation dose for use in stereotactic radiosurgery of brain or spine	Existing guideline.	http://cavernoma1.users4 Identification of optimum dose of	Change of symptoms; acceptability to	Genetic Alliance UK. Cavernoma Alliance UK.	n/a	n/a	n/a	n/a	Radiotherapy; Surgery	Neurological conditions:Brain and	Neurological conditions;		Any age
	cavernomas?		O.interdns.co.uk/wp- radiation: change in symptoms, or	patients or carers	Guidelines for the management of cerebral cavernous						conditions:Brain and	Surgery; Radiotherapy	recommendation.	
			content/uploads/2015/03/ change in management of		malformations in adults. London; Cavernoma Alliance						spinal cavernomas			
			final-CCM-guidelines.pdf symptoms; adverse effects or		UK: 2014.									
			complications; patient satisfaction; health related quality of life; service		http://cavernoma1.users40.interdns.co.uk/wp- content/uploads/2015/03/final-CCM-guidelines.pdf									
			related issues; and health related		content/uploads/2015/03/Inal-CCM-guidelines.pdf									
			related issues, and realiti related											
			cosi											
Deleted a	s merged with Q1 as agreed at the Workshop.													
Old19	Is surgical removal or no treatment better for improving outcome for people with	See uncertainty 1, above.	Prognosis: change in symptoms,							Prognosis; Surgery; Service	Neurological			
	cavernomas?		or change in management of							delivery	conditions: Brain and			
			symptoms; adverse effects or								spinal cavernomas			
			complications; patient											
			satisfaction; health related											
			quality of life; service related											
			issues; and health related cost											
Old 20	Is stereotactic radio surgery or no treatment better for improving outcome for people	See uncertainty 1, above.	Prognosis: change in symptoms,							Prognosis; Surgery; Service				
	with cavernomas?		or change in management of							delivery	conditions: Brain and			
			symptoms; adverse effects or								spinal cavernomas			
			complications; patient											
			satisfaction; health related											
			quality of life; service related											
			issues; and health related cost											
												1		