

eTable 5: Results of meta-analyses involving 14-3-3 γ and subgroup analyses based on QUADAS-2 quality and clinical criteria

Abbreviations of domains and criteria can be found in eTable 3.

FPR: false positive rate

NaN: not a number (correlation cannot be calculated)

SD: standard deviation

TPR: true positive rate

3 studies that investigated 14-3-3 γ (x: included in respective analysis)

Level of certainty of sCJD diagnosis	Studies included	SD (TPR)	SD (FPR)	Corr.	Abu-Rumeileh	Leitão	Simon
All studies							
Definite sCJD cases	2	0.58	0.99	1.00	x	x	
Definite and probable sCJD cases	3	0.42	0.85	0.61	x	x	x
Definite, probable, and possible sCJD cases	3	0.42	0.85	0.61	x	x	x
Aβ42 not investigated							
Definite sCJD cases	2	0.58	0.99	1.00	x	x	
Definite and probable sCJD cases	3	0.42	0.85	0.61	x	x	x
Definite, probable, and possible sCJD cases	3	0.42	0.85	0.61	x	x	x
Case-control studies							
Definite sCJD cases	2	0.58	0.99	1.00	x	x	
Definite and probable sCJD cases	2	0.58	0.99	1.00	x	x	
Definite, probable, and possible sCJD cases	2	0.58	0.99	1.00	x	x	
Cohort studies							
Definite sCJD cases	0						
Definite and probable sCJD cases	1						x
Definite, probable, and possible sCJD cases	1						x
D01 excl. high risk of bias (≥ 3 / 3 criteria)							
Definite sCJD cases	1					x	
Definite and probable sCJD cases	2	0.63	0.23	1.00		x	x
Definite, probable, and possible sCJD cases	2	0.63	0.23	1.00		x	x
D01 excl. high risk of bias (≥ 2 / 3 criteria)							
Definite sCJD cases	0						
Definite and probable sCJD cases	1						x

3 studies that investigated 14-3-3γ (x: included in respective analysis)

Level of certainty of sCJD diagnosis	Studies included	SD (TPR)	SD (FPR)	Corr.	Abu-Rumeileh	Leitão	Simon
Definite, probable, and possible sCJD cases D01 excl. high risk of bias (≥1 / 3 criteria)	1						x
Definite sCJD cases	0						
Definite and probable sCJD cases	1						x
Definite, probable, and possible sCJD cases D02 excl. high risk of bias (≥1 / 2 criteria)	1						x
Definite sCJD cases	0						
Definite and probable sCJD cases	1						x
Definite, probable, and possible sCJD cases D03 excl. high risk of bias (≥1 / 2 criteria)	1						x
Definite sCJD cases	2	0.58	0.99	1.00	x	x	
Definite and probable sCJD cases	2	0.58	0.99	1.00	x	x	
Definite, probable, and possible sCJD cases D04 excl. high risk of bias (≥3 / 4 criteria)	2	0.58	0.99	1.00	x	x	
Definite sCJD cases	2	0.58	0.99	1.00	x	x	
Definite and probable sCJD cases	3	0.42	0.85	0.61	x	x	x
Definite, probable, and possible sCJD cases D04 excl. high risk of bias (≥2 / 4 criteria)	3	0.42	0.85	0.61	x	x	x
Definite sCJD cases	2	0.58	0.99	1.00	x	x	
Definite and probable sCJD cases	3	0.42	0.85	0.61	x	x	x
Definite, probable, and possible sCJD cases D04 excl. high risk of bias (≥1 / 4 criteria)	3	0.42	0.85	0.61	x	x	x
Definite sCJD cases	2	0.58	0.99	1.00	x	x	
Definite and probable sCJD cases	2	0.58	0.99	1.00	x	x	
Definite, probable, and possible sCJD cases D05 excl. poor applicability (≥1 / 3 criteria)	2	0.58	0.99	1.00	x	x	
Definite sCJD cases	1				x		
Definite and probable sCJD cases	2	0	0.69	NaN	x		x
Definite, probable, and possible sCJD cases	2	0	0.69	NaN	x		x

3 studies that investigated 14-3-3γ (x: included in respective analysis)

Level of certainty of sCJD diagnosis	Studies included	SD (TPR)	SD (FPR)	Corr.	Abu-Rumeileh	Leitão	Simon
Disease duration not reported							
Definite sCJD cases	0						
Definite and probable sCJD cases	1						x
Definite, probable, and possible sCJD cases	1						x
Disease duration reported							
Definite sCJD cases	2	0.58	0.99	1.00	x	x	
Definite and probable sCJD cases	2	0.58	0.99	1.00	x	x	
Definite, probable, and possible sCJD cases	2	0.58	0.99	1.00	x	x	
K01 excl. high risk of bias							
Definite sCJD cases	0						
Definite and probable sCJD cases	1						x
Definite, probable, and possible sCJD cases	1						x
K01 excl. high/unclear risk of bias							
Definite sCJD cases	0						
Definite and probable sCJD cases	1						x
Definite, probable, and possible sCJD cases	1						x
K02 excl. high risk of bias							
Definite sCJD cases	0						
Definite and probable sCJD cases	1						x
Definite, probable, and possible sCJD cases	1						x
K02 excl. high/unclear risk of bias							
Definite sCJD cases	0						
Definite and probable sCJD cases	1						x
Definite, probable, and possible sCJD cases	1						x
K03 excl. high risk of bias							
Definite sCJD cases	1					x	
Definite and probable sCJD cases	2	0.63	0.23	1.00		x	x
Definite, probable, and possible sCJD cases	2	0.63	0.23	1.00		x	x
K03 excl. high/unclear risk of bias							

3 studies that investigated 14-3-3γ (x: included in respective analysis)

Level of certainty of sCJD diagnosis	Studies included	SD (TPR)	SD (FPR)	Corr.	Abu-Rumeileh	Leitão	Simon
Definite sCJD cases	0						
Definite and probable sCJD cases	1						x
Definite, probable, and possible sCJD cases	1						x
K04 excl. high risk of bias							
Definite sCJD cases	2	0.58	0.99	1.00	x	x	
Definite and probable sCJD cases	3	0.42	0.85	0.61	x	x	x
Definite, probable, and possible sCJD cases	3	0.42	0.85	0.61	x	x	x
K05 excl. high risk of bias							
Definite sCJD cases	0						
Definite and probable sCJD cases	1						x
Definite, probable, and possible sCJD cases	1						x
K05 excl. high/unclear risk of bias							
Definite sCJD cases	0						
Definite and probable sCJD cases	1						x
Definite, probable, and possible sCJD cases	1						x
K07 excl. high risk of bias							
Definite sCJD cases	2	0.58	0.99	1.00	x	x	
Definite and probable sCJD cases	2	0.58	0.99	1.00	x	x	
Definite, probable, and possible sCJD cases	2	0.58	0.99	1.00	x	x	
K08 excl. high risk of bias							
Definite sCJD cases	2	0.58	0.99	1.00	x	x	
Definite and probable sCJD cases	3	0.42	0.85	0.61	x	x	x
Definite, probable, and possible sCJD cases	3	0.42	0.85	0.61	x	x	x
K09 excl. high risk of bias							
Definite sCJD cases	2	0.58	0.99	1.00	x	x	
Definite and probable sCJD cases	2	0.58	0.99	1.00	x	x	
Definite, probable, and possible sCJD cases	2	0.58	0.99	1.00	x	x	
K09 excl. high/unclear risk of bias							
Definite sCJD cases	2	0.58	0.99	1.00	x	x	

3 studies that investigated 14-3-3γ (x: included in respective analysis)

Level of certainty of sCJD diagnosis	Studies included	SD (TPR)	SD (FPR)	Corr.	Abu-Rumeileh	Leitão	Simon
Definite and probable sCJD cases	2	0.58	0.99	1.00	x	x	
Definite, probable, and possible sCJD cases	2	0.58	0.99	1.00	x	x	
K10 excl. high risk of bias							
Definite sCJD cases	2	0.58	0.99	1.00	x	x	
Definite and probable sCJD cases	3	0.42	0.85	0.61	x	x	x
Definite, probable, and possible sCJD cases	3	0.42	0.85	0.61	x	x	x
K10 excl. high/unclear risk of bias							
Definite sCJD cases	2	0.58	0.99	1.00	x	x	
Definite and probable sCJD cases	3	0.42	0.85	0.61	x	x	x
Definite, probable, and possible sCJD cases	3	0.42	0.85	0.61	x	x	x
K11 excl. high/unclear risk of bias							
Definite sCJD cases	2	0.58	0.99	1.00	x	x	
Definite and probable sCJD cases	3	0.42	0.85	0.61	x	x	x
Definite, probable, and possible sCJD cases	3	0.42	0.85	0.61	x	x	x
K12 excl. poor applicability							
Definite sCJD cases	1				x		
Definite and probable sCJD cases	2	0	0.69	NaN	x		x
Definite, probable, and possible sCJD cases	2	0	0.69	NaN	x		x
K12 excl. poor/unclear applicability							
Definite sCJD cases	1				x		
Definite and probable sCJD cases	1				x		
Definite, probable, and possible sCJD cases	1				x		
K13 excl. poor applicability							
Definite sCJD cases	2	0.58	0.99	1.00	x	x	
Definite and probable sCJD cases	3	0.42	0.85	0.61	x	x	x
Definite, probable, and possible sCJD cases	3	0.42	0.85	0.61	x	x	x
K13 excl. poor/unclear applicability							
Definite sCJD cases	2	0.58	0.99	1.00	x	x	
Definite and probable sCJD cases	3	0.42	0.85	0.61	x	x	x

3 studies that investigated 14-3-3 γ (x: included in respective analysis)

Level of certainty of sCJD diagnosis	Studies included	SD (TPR)	SD (FPR)	Corr.	Abu-Rumeileh	Leitão	Simon
Definite, probable, and possible sCJD cases	3	0.42	0.85	0.61	x	x	x
Male-to-female ratio <0.75							
Definite sCJD cases	1				x		
Definite and probable sCJD cases	1				x		
Definite, probable, and possible sCJD cases	1				x		
Male-to-female ratio >1							
Definite sCJD cases	0						
Definite and probable sCJD cases	1						x
Definite, probable, and possible sCJD cases	1						x
Male-to-female ratio 0.75–1							
Definite sCJD cases	1					x	
Definite and probable sCJD cases	1					x	
Definite, probable, and possible sCJD cases	1					x	
Median age >65 years							
Definite sCJD cases	1				x		
Definite and probable sCJD cases	2	0	0.69	NaN	x		x
Definite, probable, and possible sCJD cases	2	0	0.69	NaN	x		x
Not only 14-3-3β investigated							
Definite sCJD cases	1					x	
Definite and probable sCJD cases	1					x	
Definite, probable, and possible sCJD cases	1					x	
RT-QuIC not investigated							
Definite sCJD cases	1					x	
Definite and probable sCJD cases	1					x	
Definite, probable, and possible sCJD cases	1					x	
Studies >2009							
Definite sCJD cases	2	0.58	0.99	1.00	x	x	
Definite and probable sCJD cases	3	0.42	0.85	0.61	x	x	x
Definite, probable, and possible sCJD cases	3	0.42	0.85	0.61	x	x	x

3 studies that investigated 14-3-3γ (x: included in respective analysis)

Level of certainty of sCJD diagnosis	Studies included	SD (TPR)	SD (FPR)	Corr.	Abu-Rumeileh	Leitão	Simon
Studies with a priori cut-off							
Definite sCJD cases	1					x	
Definite and probable sCJD cases	2	0.63	0.23	1.00		x	x
Definite, probable, and possible sCJD cases	2	0.63	0.23	1.00		x	x
Studies with data-driven cut-off							
Definite sCJD cases	1				x		
Definite and probable sCJD cases	1				x		
Definite, probable, and possible sCJD cases	1				x		
Study sample: only definite cases							
Definite sCJD cases	2	0.58	0.99	1.00	x	x	
Definite and probable sCJD cases	3	0.42	0.85	0.61	x	x	x
Definite, probable, and possible sCJD cases	3	0.42	0.85	0.61	x	x	x