

The prevalence of *EGFR* mutation in patients with non-small cell lung cancer: a systematic review and meta-analysis

APPENDIX TABLES

Table A.1: The basic characteristics of included studies

See Appendix File 1

Table A.2: Post-hoc subgroup analysis on the detection methods, samples for EGFR mutation, and publication year

Group variables	No. of studies	Mutation prevalence, 95% CI (%)	Tests of heterogeneity	
			P	I ² (%)
All studies	456	32.3 (30.9 to 33.7)	<0.001	97.3
Detection method				
Pyrosequencing or direct sequencing	306	32.2 (30.3 to 34.0)	<0.001	97.4
COLD-PCR and pyrosequencing or fragment size analysis	7	31.9 (24.5 to 39.3)	<0.001	85.8
Sanger sequencing	2	35.5 (27.2 to 43.7)	0.205	37.8
Amplification refractory mutation system ARMS	28	33.7 (27.1 to 40.4)	<0.001	97.8
Real-time PCR	13	27.6 (20.2 to 35.1)	<0.001	96.9
Fragment size analysis	4	42.5 (29.0 to 56.1)	<0.001	89.5
RT-PCR	7	41.6 (23.6 to 59.5)	<0.001	97.0
PCR and sequencing	26	28.9 (24.9 to 32.9)	<0.001	95.7
PCR single strand conformational polymorphism	3	43.0 (0.0 to 93.1)	<0.001	99.5
High Performance Liquid Chromatography	11	37.7 (32.7 to 42.7)	<0.001	70.8
PCR clamp method or PCR invader assay	5	52.9 (36.2 to 69.7)	<0.001	93.3
Peptide nucleic acid (PNA) clamping	21	33.4 (28.1 to 38.8)	<0.001	91.3
Other methods	2	14.1 (10.4 to 17.7)	0.931	0.0
Unclear	21	25.0 (20.9 to 29.1)	<0.001	96.2
Sample for <i>EGFR</i> mutation detection				
Tissue	365	32.8 (31.1 to 34.5)	<0.001	97.0
Blood	10	36.0 (26.4 to 45.6)	<0.001	88.3
Mixed (at least include tissue or blood samples)	41	31.9 (26.8 to 37.0)	<0.001	97.5
Other samples	11	25.1 (14.2 to 36.1)	<0.001	94.9
Unclear	29	27.8 (24.1 to 31.5)	<0.001	97.7
Publication date				
2004	5	38.3 (20.9 to 55.7)	<0.001	96.0
2005	36	26.6 (22.0 to 31.2)	<0.001	94.8
2006	29	29.2 (24.7 to 33.7)	<0.001	87.3
2007	38	30.6 (24.8 to 36.4)	<0.001	96.5
2008	37	34.2 (28.3 to 40.1)	<0.001	95.8
2009	37	30.0 (25.0 to 35.1)	<0.001	96.6
2010	60	35.3 (31.0 to 39.6)	<0.001	96.9
2011	107	34.9 (31.5 to 38.4)	<0.001	97.9
2012	89	31.3 (28.5 to 34.2)	<0.001	97.9
2013	18	31.2 (24.1 to 38.3)	<0.001	97.6

Table A.3: The pooled prevalence of *EGFR* mutation in studies from China

Group variables	No. of studies	Mutation prevalence, 95%CI (%)	Tests of heterogeneity	
			P	I ² (%)
All studies	104	38.4(35.7 to 41.1)	<0.001	93.0
Exons				
Exon 19	86	19.9(18.0 to 21.9)	<0.001	88.8
Exon 21	84	15.8(14.3 to 17.4)	<0.001	83.2
Exon 19 or 21	34	35.8(31.4 to 40.2)	<0.001	91.9
Gender				
Female	82	50.3(46.9 to 53.7)	<0.001	84.8
Male	82	30.3(27.5 to 33.2)	<0.001	88.0
Smoking status				
Non-smoker	69	49.0(45.3 to 52.7)	<0.001	89.4
Past or current smoker	65	27.6(24.2 to 31.0)	<0.001	90.0
Histology				
Adenocarcinoma	82	50.3(46.9 to 53.7)	<0.001	84.8
Non-adenocarcinoma	60	16.9(14.3 to 19.5)	<0.001	81.4
Chemotherapy				
Chemotherapy	15	37.4(29.0 to 45.9)	<0.001	90.2
No chemotherapy	25	36.3(30.3 to 42.3)	<0.001	93.4

Table A.4: The Pooled prevalence of *EGFR* mutation in studies from Japan

Group variables	No. of studies	Mutation prevalence, 95% CI (%)	Tests of heterogeneity	
			P	I ² (%)
All studies	107	36.6(33.2 to 40.0)	<0.001	96.4
Exons				
Exon 19	81	17.8(15.9 to 19.7)	<0.001	87.8
Exon 21	79	15.9(14.2 to 17.5)	<0.001	83.2
Exon 19 or 21	27	35.3(30.6 to 39.9)	<0.001	93.1
Gender				
Female	81	49.8(45.3 to 54.2)	<0.001	91.8
Male	81	22.5(19.7 to 25.3)	<0.001	91.5
Smoking status				
Non-smoker	66	54.4(48.6 to 60.1)	<0.001	94.7
Past or current smoker	67	20.6(17.9 to 23.4)	<0.001	90.8
Histology				
Adenocarcinoma	76	41.6(37.2 to 46.0)	<0.001	95.7
Non-adenocarcinoma	42	3.3(2.4 to 4.1)	0.068	25.8
Chemotherapy				
Chemotherapy	5	45.3(37.7 to 52.8)	0.672	0.0
No chemotherapy	17	36.5(27.3 to 45.8)	<0.001	92.1

Table A.5: The pooled prevalence of *EGFR* mutation in studies from Korea

Group variables	No. of studies	Mutation prevalence, 95% CI (%)	Tests of heterogeneity	
			P	I ² (%)
All studies	48	32.4(28.0 to 36.8)	<0.001	94.8
Exons				
Exon 19	37	17.8(15.0 to 20.6)	<0.001	88.4
Exon 21	37	10.2(8.3 to 12.1)	<0.001	82.9
Exon 19 or 21	4	37.9(25.4 to 50.5)	<0.001	89.9
Gender				
Female	38	47.5(43.2 to 51.7)	<0.001	80.5
Male	35	24.7(20.2 to 29.2)	<0.001	90.7
Histology				
Adenocarcinoma	33	40.6(36.2 to 45.0)	<0.001	89.2
Non-adenocarcinoma	25	14.4(10.5 to 18.3)	<0.001	81.4
Smoking status				
Non-smoker	35	47.2(42.9 to 51.5)	<0.001	80.0
Past or current smoker	35	23.9(19.7 to 28.2)	<0.001	88.6
Chemotherapy				
Chemotherapy	8	34.8(18.4 to 51.2)	<0.001	97.2
No chemotherapy	5	19.8(16.4 to 23.3)	0.878	0.0

Table A.6: The pooled prevalence of *EGFR* mutation in studies from U.S.

Group variables	No. of studies	Mutation prevalence, 95% CI (%)	Tests of heterogeneity	
			P	I ² (%)
All studies	68	23.9(21.3 to 26.5)	<0.001	96.6
Exons				
Exon 19	37	12.8(11.0 to 14.6)	<0.001	92.9
Exon 21	34	7.6(6.3 to 9.0)	<0.001	92.0
Exon 19 or 21	19	25.0(21.5 to 28.5)	<0.001	92.2
Gender				
Female	42	29.3(25.4 to 33.2)	<0.001	93.4
Male	38	18.9(15.7 to 22.2)	<0.001	89.7
Smoking status				
Non-smoker	42	46.5(41.3 to 51.6)	<0.001	91.8
Past or current smoker	40	14.3(12.5 to 16.1)	<0.001	83.2
Histology				
Adenocarcinoma	42	25.3(22.7 to 27.9)	<0.001	90.1
Non-adenocarcinoma	21	17.3(12.7 to 21.9)	<0.001	79.6
Chemotherapy				
Chemotherapy	5	19.1(6.2 to 31.9)	<0.001	96.9
No chemotherapy	7	19.8(13.5 to 26.2)	<0.001	92.8

Table A.7: Search strategy

Search	Query
1	EGFR
2	lung cancer
3	1 and 2
4	mutation or mutant or mutated
5	3 and 4

LIST OF ELIGIBLE STUDIES

- Fan M, Li H, Lü Z, Zhang H, Shen X. [Relationship of epidermal growth factor receptor gene mutations, clinicopathologic features and prognosis in patients with non-small cell lung cancer]. *Chinese journal of pathology*. 2011; 40:679-682.
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