

**Supplementary Table S1.** Basic characteristics of studies on the prevalence of HHcy (defined as Hcy  $\geq$  15 $\mu$ mol/L) in China

Authors	Year of Publication	year of data collection	Population source	Age (Years)	Province	Setting	Area	Study Location	Methods of Hcy measurement	Sample	Number of Cases	Prevalence of HHcy (%)	Prevalence of male (%)	Prevalence of female (%)
Pang2014[1]	2014	2011.5-2012.11	Community-based	42.2 $\pm$ 11.4	Xinjiang	R	N	I	ECA	1003	316	31.5	49.9	17.5
Wang2014[2]	2014	NR	Community-based	54.05 $\pm$ 12.11	Guangdong	U	S	Co	ECA	1586	270	17.02	29.6	8.47
Liu2014[3]	2014	2012.08-2013.09	Physical examination - based	43.04 $\pm$ 10.24	Jilin	U and R	N	I	ECA	1106	200	18.1	NR	NR
Liu2015[4]	2015	2007	Community-based	43.57	Shanxi	U and R	N	I	ECA	2645	1791	67.7	81.4	58.8
Zhang2015[5]	2015	2009	Community-based	69.5 $\pm$ 8.1	Beijing	U	N	I	ECA	1458	902	61.87	NR	NR
Deng2015[6]	2015	NR	Community-based	72.96 $\pm$ 7.66	Shanghai	R	C	Co	HPLC	1621	569	35.1	43.6	29.5
Wang2016[7]	2016	2013.1-2013.12	Physical examination - based	49.1 $\pm$ 11.7	Guangdong	U and R	S	Co	IA	1136	575	50.61	65.61	27.78
Wu2016a[8]	2016	2013.1-2014.10	Physical examination - based	45.86 $\pm$ 13.53	Sichuan	U and R	C	I	ECA	11740	3393	28.9	40.53	8.77
Wu2016b[9]	2016	2014.1	Physical examination - based	38.0 $\pm$ 8.6	Shanghai	U and R	C	Co	ECA	438	125	28.54	NR	NR
Zhang2016[10]	2016	2012.8-2012.12	Community-based	64.6 $\pm$ 7.4	Shanxi	R	N	I	ECA	2355	1710	72.6	84.3	63.2
Chen2017[11]	2017	2012.7-2013.8	Community-based	53.8 $\pm$ 10.6	Liaoning	R	N	I	ECA	7365	2944	39.97	59.04	25.79
Zhan2017[12]	2017	2012.8-2012.12	Community-based	63.68	Shanxi	U	N	I	ECA	516	250	48.4	NR	NR
Wei2017[13]	2017	2015-2016	Community-based	$\geq$ 35	Guangxi	R	S	Co	ECA	600	214	35.7	40.4	28.1

Author	Year	Study Period	Method	Prevalence	Region	Design	Setting	Control	Exposure	Outcome	Number of Cases	Number of Controls	OR	95% CI	Number of Controls
Yang2018[14]	2018	2013-2018	Physical examination - based	54.8±7.9	Hunan	U and R	C	Co	NR	207069	82701	39.94	55.6	16.7	
Qin2018[15]	2018	2015-2016	Physical examination - based	≥20	Guangxi	U and R	S	Co	ECA	8043	4086	50.8	52.3	48.1	
Chen2018[16]	2020	2016.08-2016.09	Community-based	61.2±9.5	Jiangsu	U	C	Co	ECA	16644	5018	30.15	43.65	22.31	
Wang2019[17]	2019	2013	Community-based	55.7 ± 11.1	Shanghai	U and R	C	Co	ECA	8337	2059	24.7	NR	NR	
Cao2019[18]	2019	2015.07-2017.12	Physical examination - based	NR	Chongqing	U and R	C	I	ECA	7308	1237	16.93	90.54	5.6	
Jiang2019[19]	2019	2007	Community-based	35-64	Jiangxi	R	C	I	ECA	3886	1542	39.68	49.61	24.14	
Tan2019[20]	2019	2015	Physical examination - based	72.57±6.51	Shanghai	U and R	C	Co	ECA	5976	2487	41.62	51.47	33.82	
Zhang2019[21]	2019	NR	Physical examination - based	61.91±7.26	Heilongjiang	U	N	I	ELISA	668	200	29.94	25.99	33.72	
Zhao2019[22]	2019	2017.01	Physical examination - based	59.8±9.8	Xinjiang	U and R	N	I	NR	7492	4745	63.33	74.23	54.46	
Deng2020[23]	2020	2017.07-2018.10	Physical examination - based	50.37±6.09	Guangxi	U and R	S	Co	ECA	1768	270	15.2	23.1	6	
Wang2020[24]	2020	2018.01-2018.06	Physical examination - based	58.15±11.74	Hebei	U	N	I	NR	2572	724	28.15	NR	NR	
Yu2020[25]	2020	2001.01-2003.08	Community-based	54 ± 10	Liaoning	R	N	Co	ECA	6837	2824	41.3	NR	NR	
Cui2020[26]	2020	2016.06-2017.11	Community-based	20-74	Shanghai	U	C	Co	ECA	8201	2673	32.59	52.69	18.6	
Niu2021[27]	2021	2013-2014	Community-based	54.5	Hunan	U and R	C	Co	NR	4012	1420	35.39	45.38	28.46	

Guo2021[28]	2021	2012.03-2015.07	Community-based	57.5±10.9	Guangdong	U	S	Co	NR	15808	2739	17.33	NR	NR
Liu2021[29]	2021	2014.04-2015.04	Physical examination - based	40±8.4	Sichuan	U and R	C	I	ELISA	470	163	34.68	NR	NR

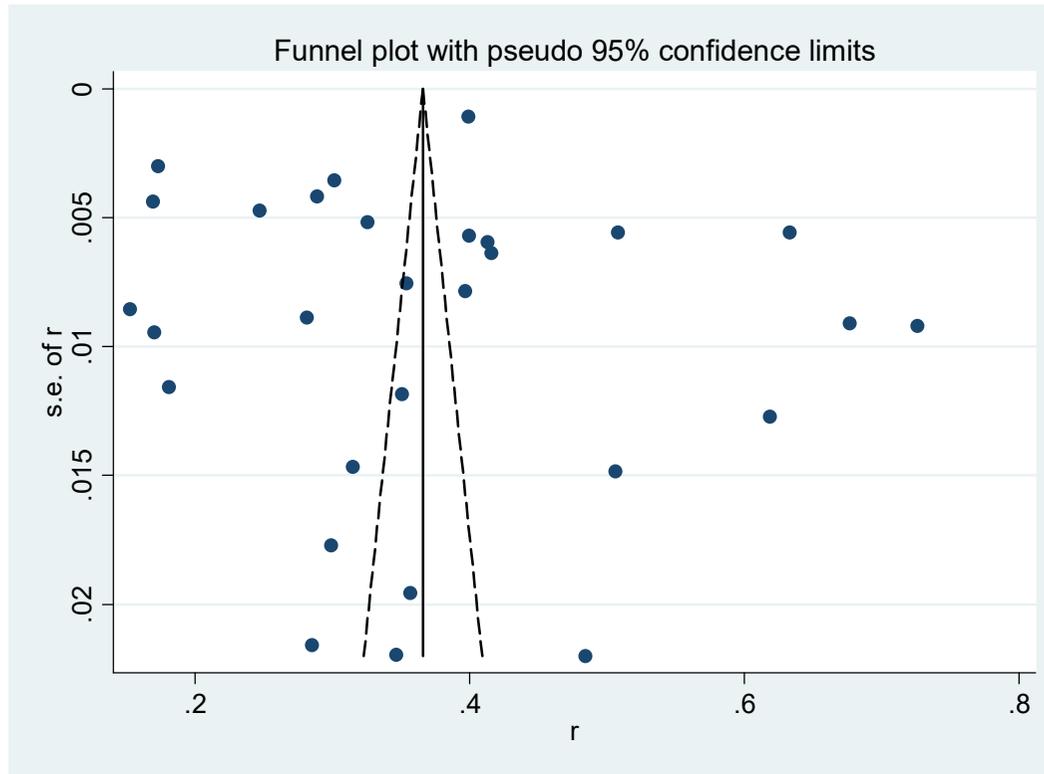
HHcy, hyperhomocysteinemia; Hcy, homocysteine; NR, not reported; N, north; S, south; Ce, central; I, inland; Co, coastal; U, urban; R, rural; HPLC, high-performance liquid chromatography; ECA, enzymatic cycling assay; IA, immunoturbidimetric assays.

**Table S2. Quality scores for assessing the risk of bias of in the included articles for observation study.**

Study(year)	Q1	Q2	Q3	Q4	Q5	Q6	Q7	Q8	Q9	Q10	Q11	Total Score
Pang2014[1]	1	0	1	1	1	0	1	1	0	1	1	8
Wang2014[2]	1	1	1	1	1	0	1	0	0	1	1	8
Liu2014[3]	1	0	1	1	1	0	0	0	0	0	1	5
Liu2015[4]	1	1	1	1	1	0	1	1	0	1	1	9
Zhang2015[5]	1	1	1	1	1	0	1	1	0	1	1	9
Deng2015[6]	1	0	0	1	1	0	0	0	0	1	1	5
Wang2016[7]	1	1	1	1	1	0	0	0	0	0	1	6
Wu2016a[8]	1	0	1	1	1	0	0	0	0	0	1	5
Wu2016b[9]	1	1	1	1	1	0	1	0	0	1	1	8
Zhang2016[10]	1	1	1	1	1	0	1	0	0	1	1	8
Chen2017[11]	1	1	1	1	1	0	0	1	0	1	1	8
Zhan2017[12]	1	1	1	1	1	0	1	1	0	1	1	9
Wei2017[13]	1	1	1	1	1	0	0	0	0	0	1	6
Yang2018[14]	1	1	1	1	1	0	0	0	0	0	1	6
Qin2018[15]	1	1	1	1	1	0	0	1	0	0	1	7

Chen2018[16]	1	1	1	1	1	0	0	0	0	0	1	6
Wang2019[17]	1	1	1	1	1	0	0	1	0	0	1	7
Cao2019[18]	1	1	1	1	1	0	0	1	0	0	1	7
Jiang2019[19]	1	0	1	1	1	0	0	0	0	0	1	5
Tan2019[20]	1	0	1	1	1	0	1	0	0	1	1	7
Zhang2019[21]	1	1	0	1	1	0	1	0	0	1	1	7
Zhao2019[22]	1	1	1	1	1	0	0	0	0	0	1	6
Deng2020[23]	1	1	1	1	1	0	0	1	0	0	1	7
Wang2020[24]	1	1	1	1	1	0	1	0	0	1	1	8
Yu2020[25]	1	1	1	1	1	0	1	1	0	1	1	9
Cui2020[26]	1	1	1	1	1	0	1	1	0	1	1	9
Niu2020[27]	1	1	1	1	1	0	1	1	0	1	1	9
Guo2021[28]	1	1	1	1	1	0	0	1	0	1	1	8
Liu2021[29]	1	1	1	1	1	0	0	0	0	0	1	6

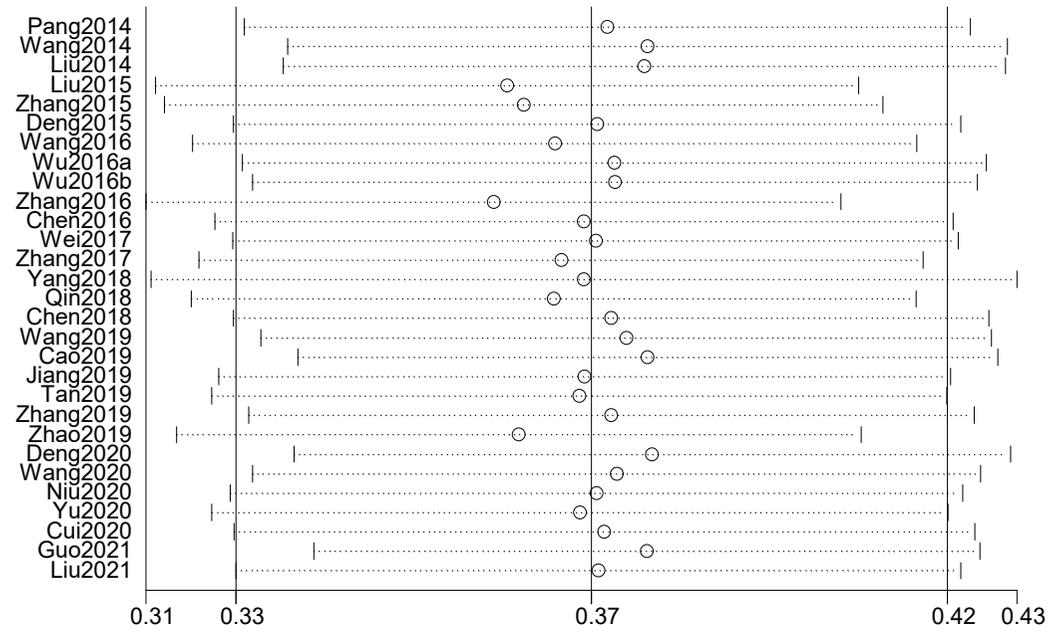
“No” or “Unclear” was scored “0”, and “Yes” was scored “1”. Low quality:  $\leq 3$  scores; Moderate quality: 4-7 scores; High quality:  $\geq 8$  scores. Q1: Define the source of information (survey, record review); Q2: List inclusion and exclusion criteria for exposed and unexposed subjects (cases and controls) or refer to previous publications; Q3: Indicate time period used for identifying patients; Q4: Indicate whether or not subjects were consecutive if not population-based; Q5: Indicate if evaluators of subjective components of study were masked to other aspects of status of the participants; Q6: Describe any assessments undertaken for quality assurance purpose; Q7: Explain any patient exclusions from analysis; Q8: Describe how confounding was assessed and/or controlled; Q9: If applicable, explain how missing data were handled in the analysis; Q10: Summarize patient response rates and completeness of data collection; Q11: Clarify what follow-up, if any, was expected and the percentage of patients for which incomplete data or follow-up was obtained.



Supplementary Figure S1. Funnel chart

Meta-analysis random-effects estimates (linear form)

Study omitted



Supplementary Figure S2. Sensitivity analyses

1. Pang,R.H.; Zhang,J.Y.; Xu,S.Z.; Ma,R.L.; Guo,H.; Liu,J.M.; Zhang,M.; Ding,Y.S.; Li,S.G.; Guo,S,X. The Epidemiological Characteristics of High Serum Level of Homocysteine in Kazakh Population at Xinyuan County of Xinjiang. *Chin Circ J.* 2014, 29, 913-916.
2. Wang,C.Y.; Zhang,T.; Chen,S.H.; Ma,J.P.; Liu,S.Y.; Peng,X.L.; Peng,J.; Xu,J.; Chen,Z.W. Study on the distribution of hyperhomocysteinemia and its influencing factors in healthy population of community in Nanshan Shenzhen. *J Trop Med.* 2014, 14, 1204-1208.
3. Liu, .R.; Jiang, W.X.; Song, C.X.; Zheng, W.M.; Tan L. Hyperhomocysteinemia and its associated risk factors in healthy subjects. *Chinese journal of clinical research.* 2014. 27. 637-638.
4. Liu, X.D.; Gao, B.; Sun, D.; Shi, M.;Ma, Y.Y.; Liu, Z.R.; Wang, B.; Xu, X.; Xu, X.; Ji, Q.H.; Zhao, G. Prevalence of hyperhomocysteinaemia and some of its major determinants in Shaanxi Province, China: a cross-sectional study. *Br J Nutr.* 2015, 113, 691-698. <http://doi.org/10.1017/s0007114514004218>
5. Zhang,Z.Y.;Fang,X.H.;Jie,X.H.;Tang,Z.;Guan,S.C.;Liu,H.J.;Wu,X.G.;Wang,C.X.;Hou,C.B.;Gu,X.The association between serum homocysteine and blood pressure among middle-aged and elderly people. *Chin J hypertens.* 2015, 23, 846-850.
6. Deng,Q.Q.;Gao,L.L.;Fu,J.H.; Pan,D.H.; Xiao,W.Z.; Li,Q.H.; Ding,D.; Pan,X.J.; Yu,B. The prevalence and risk factors of hyperhomocysteinemia in elderly people in rural community of Shanghai. *Fudan Univ J Med Sci.* 2015, 42, 722-726.
7. Wang,J.F.; Ma,X.T.; Liu,G.Z.; Xie,T.; Xie,Y.L. The characteristic of total plasma total homocysteine level distribution in population of Zhaihai area. *Chongqing Med J.* 2016, 45, 3972-3974.xx
8. Wu,Q.Q.; Tang,H.R.; Tang,G.H.; Li,Q.; Xue,G.C.; Chen,Y.; Leng,R.;Wang,Y.J. The Cross-sectional Survey of the Serum Homocysteine in Chengdu Area. *Sichuan Med J.* 2016, 37, 783-787.
9. Wu,Y,Z; Lu,D.R.; Chen,X.X.; Jiang,Y.M. Correlation of homocysteine levels with gene polymorphisms of metabolic enzyme in 438 adults taking physical examination in Shanghai,china.*Acad J Second Military Med Univ.* 2016, 37, 936-942.
10. Zhang, J.;Liu, T.T.;Zhang, W.;Li, Y.;Niu, X.Y.;Fang, Y.L.;Ma, L.S.;Li, C.X. Hyperhomocysteinemia Is Associated with Vitamin B-12 Deficiency: A Cross-sectional Study in a Rural, Elderly Population of Shanxi China. *J Nutr Health Aging.* 2016, 20, 594-601. <http://doi.org/10.1007/s12603-015-0650-511>
11. Chen, S.;Guo, X.;Dong, S.;Li, Z.;Sun, Y. Relationship between lifestyle factors and hyperhomocysteinemia in general Chinese population: a cross-sectional study. *Postgrad Med.* 2017, 129, 216-223. <http://doi.org/10.1080/00325481.2017.1255126>
12. Zhang, H.; Shang, S.H.; Deng, M.Y.; Yan, F.; Li, Z.L.,; Chen, C.; Jiang, Y.; Dang, L.J.; Zhu, J.; Li, Y.; Qu, Q.M. Correlation between plasma homocysteine level and blood pressure in the high-risk population of stroke in the community of Xi'an. *Chinese Journal of Integrative Medicine on Cardio-Cerebrovascular Disease.* 2017,15, 3186-3193.

13. Wei,S.W.; Wei,Y.L.; Xun,W.H.; Wei,X.Y. Prevalence and Influencing Factors of Hyperhomocysteinemia in Baiku Yao Nationality Residents in Guangxi. *PJCCPVD*. 2017, 25, 73-75.
14. Yang,B.F. Correlation analysis between overweight and obesity and plasma homocysteine and blood lipid levels in healthy people. *Chin commun doc*. 2018, 34, 122-123.
15. Qin, Y.Y.;Wang, P.;Qin, J.Q.;Wei, A.Q.;Huang, P.;Lai, Z.F.;Lin, F.Q. Prevalence of hyperhomocysteinemia during routine physical examination in Guangxi Province, China and related risk factors. *J Clin Lab Anal*. 2018, 32. <http://doi.org/10.1002/jcla.22178>
16. Chen, L.;Wang, B.;Wang, J.;Ban, Q.;Wu, H.;Song, Y.;Zhang, J.;Cao, J.;Zhou, Z.;Liu, L.;Cao, T.;Gao, L.;Guo, H.;Zhang, T.;Tang, G.;Huang, X.;Zhang, Y.;Li, J.;Huo, Y.;Cheng, X.;Zang, T.;Xu, X.;Zhang, H.;Qin, X. Association between serum total homocysteine and arterial stiffness in adults: a community-based study. *J Clin Hypertens (Greenwich)*. 2018, 20, 686-693. <http://doi.org/10.1111/jch.13246>
17. Wang, Y.H.;Yan, Q.H.;Xu, J.Y.;Li, X.J.;Cheng, M.N. High Prevalence and Factors Contributing to Hyperhomocysteinemia, Folate Deficiency, and Vitamin B12 Deficiency among Healthy Adults in Shanghai, China. *Biomed Environ Sci*. 2019, 32, 63-67. <http://doi.org/10.3967/bes2019.010>
18. Cao,Y.;Li,B.;He,J.T.; Yue,Y.; Wang,X.H.; Chen,Y.; Jia,S.Y. The relationship between serum homocysteine level and changes of cerebral hemodynamic indexes. *Chongqing Med J*. 2019, 48, 1225-1227+1230.
19. Jiang,L.X.; Dong,M.H.; Shi,S.H.; Zhang,H.J.; Li,S.M.; Xia,Y; Cao,J. The study of environmental factors for hyperhomocysteinemia in the population of Ganzhou. *J Gannan Med Univ*. 2019, 39, 762-765.
20. Tan,Q.; Wang,J.G.; Li,Y.; Wang,D.; Chen,Y.; Guo,H.Q. Clinical Characteristics of Hyperhomocysteinemia at Various Diagnostic Cut-off Thresholds. *Prevention and Treatment of Cardio-Cerebral-Vascular Disease*. 2019, 19, 292-295.
21. Zhang,Y.; Ying,X.;Guo,Y.p.; Chen,Y.L.; Wang,J.P.; Wang,Z.G. Analysis of influencing factors of homocysteine in middle-aged and elderly population. *J Harbin Med univ*. 2019, 53, 598-601.
22. Zhao,Y.; Shang,L.X.; Zhang,W.H.; Li,Y.D.; Zhou,X.H.;Tang,B.P. Correlation between metabolic syndrome components and hyperhomocysteinemia in residents with physical examination of Xinjiang area. *Chin J Prevent Control Chronic Dis*. 2019, 27, 425-427+431.
23. Deng,Z.K.; Ming,Y.; Li,H.B.; Lin,H.L.; Yang,J.; Bo,F. Analysis on the relationship between hyperhomocysteinemia and blood lipid and BMI in healthy people taking medical examination in Guilin . *Chinese Journal of Healthcare and Medicine*. 2020, 22, 599-602.
24. Wang,Z.L.; Zhang,X.M.; Hao,Z.H.; Nie,Q.; Liu,C.H.; Liu,X.X.;Wang,L.L. Relationship between serum homocysteine and metabolic syndrome in middle-aged and aged people of state-run institutions. *Pract Geriatr*. 2020, 34, 1165-1169.
25. Yu, S.;Chen, Y.;Yang, H.;Guo, X.;Zheng, L.;Sun, Y. Hyperhomocysteinemia accompany with metabolic syndrome increase the risk of left ventricular hypertrophy in

rural Chinese. *BMC Cardiovasc Disord.* 2020, 20, 44. <http://doi.org/10.1186/s12872-020-01350-2>

26. Cui,S.H.; Zhao,Q.; Wang,N.; Zhang,Y.; Qiu,Y.; Zhou,X.Y.; Yu,Y.T.; Wang,R.P.; Jiang,Y.G.; Zhao,G.M. Association between metabolic syndrome and its components with hyperhomocysteinemia in community population. *Chin J Dis Control & Prev.* 2020, 24, 1003-1008.
27. Niu, X.;Chen, J.;Wang, J.;Li, J.;Zeng, D.;Wang, S.;Hong, X. A Cross-sectional Study on the Relationship Between Homocysteine and Lipid Profiles Among Chinese Population from Hunan. *Lipids.* 2021, 56, 93-100. <http://doi.org/10.1002/lipd.12279>
28. Guo,Z.G.; Huang,J.Y.; Meng,R.W.; Shi,O.M.; Tie,X.H.; Mo,H.M.; Wang,Z.H. Correlation between high homocysteine and the risk of stroke in community residents aged 40 years and above in Shenzhen city. *Chin J Health Manage .* 2021, 15, 37-43. <http://doi.org/10.3760/cma.j.cn115624-20200527-00427>
29. Liu,X.Y.;Li,Y.;Liu,T. Correlation of homocysteine levels with gene polymorphisms of homocysteine metabolic enzymes in physical examination in Nan Chong. *J of Pub Health and Prev Med.* 2021, 32, 113-116.