

Appendix 2: Supplementary tables [posted as supplied by author]

Table A. Study characteristics of all the studies included in meta-analysis

Study	Country/ Region	Prediabetes definition and prevalence (%)	Sample size (% women)	Age (y), average (range or SD)	Follow-up (y)	Baseline CVD excluded	Events for analysis	Possibility of enrolling patients with diabetes
Stengard 1992 ²⁸	Finland	IGT (36.7%)	637 (0%)	72.4 (65-84)	5	No	CVD mortality All-cause mortality	Yes
Barzilay 1999 ²⁹	United States	IFG-WHO (14.6%) IGT (32.1%)	4515 (58%)	73 (5.6)	8	Yes	CVD incidence	No
DECODE 2001 ³⁰	Europe	IFG-WHO (10.1%) IGT (11.4%)	22514 (31.7%)	53 (30-89)	8.8	No	CVD mortality CHD mortality All-cause mortality	No
Mazza 2001 ³¹	Italy	IFG-WHO (NA)	3282 (61.0%)	73.8 (≥65)	14	No	Stroke mortality	Yes
Saydah 2001 ³²	United States	IFG-WHO (NA) IGT (17.0%)	3092 (54.3%)	49.6 (30-74)	13.5	No	CVD mortality All-cause mortality	No
Henry 2002 ³³	France	IFG-WHO (17.0%)	63443 (0%)	NA (21-60)	8	Yes	CVD mortality All-cause mortality	Yes
Rodriguez 2002 ³⁴	Japan	IFG-WHO (22.4%) IGT (26.7%)	2034 (0%)	NA (71-93)	7	Yes	CVD mortality All-cause mortality	No
Lu 2003 ³⁵	United States	IFG-WHO (10.8%)	4304 (56.1%)	56.1 (45-74)	9	No	CVD mortality All-cause mortality	Yes
Hunt 2004 ³⁶	United States	IFG-WHO (2.2 %)	2815 (56.9%)	43.4 (25-46)	12.7	Yes	CVD mortality All-cause mortality	Yes
Nakagami 2004 ³⁷	Asia	IFG-WHO (6.8%) IGT (16.2%)	6817 (54.2%)	50 (30-89)	5	No	CVD mortality All-cause mortality	No
Nakanishi 2004 ³⁸	Japan	IFG-WHO (4.0%)	6182 (0%)	47.5 (35-59)	7	Yes	CVD incidence	Yes
Tai 2004 ³⁹	Singapore	IFG-ADA (18.3%)	5091 (NA)	37.2 (12)	9.1	Yes	CHD incidence	No
Tai 2004 ³⁹	Singapore	IGT (13.1%)	3568 (NA)	NA	8	Yes	CVD incidence CHD incidence	No
Hiltunen 2005 ⁴⁰	Finland	IGT (34.0%)	379 (62.8%)	76 (NA)	9.8	No	All-cause mortality	No

Study	Country/ Region	Prediabetes definition and prevalence (%)	Sample size (% women)	Age (y), average (range or SD)	Follow-up (y)	Baseline CVD excluded	Events for analysis	Possibility of enrolling patients with diabetes
McNeill 2005 ⁴¹	United States	IFG-WHO (12.3%)	12089 (56.9%)	54 (45-64)	11	Yes	CHD incidence	Yes
Wild 2005 ⁴²	UK	IFG-WHO (11.9%) IGT (8.3%)	1496 (49.3%)	64.8 (55-74)	12.6	No	CVD mortality All-cause mortality	No
Kaarisalo 2006 ⁴³	Finland	IGT (12.3%)	1032 (52.1%)	70 (NA)	9.6	No	Ischaemic stroke	No
McNeill 2006 ⁴⁴	United States	IFG-ADA (45.5%) IFG-WHO (16.5%)	3585 (62%)	72 (65-92)	11	Yes	CHD incidence	Yes
Palmieri 2006 ⁴⁵	Italy	IFG-WHO (7.7%)	20447 (63.6%)	50.4 (35-69)	10.4	No	CHD incidence	Yes
Barr 2007 ⁴⁶	Australia	IFG-WHO (5.8%) IGT (12.4%)	10428 (54.8%)	51.4 (≥25)	5.2	No	CVD mortality All-cause mortality	No
Liu 2007 ⁸	China	IFG-ADA (21.1%)	30378(46.5%)	46.9 (35-64)	10	Yes	CVD incidence CHD incidence Stroke incidence	Yes
Nilsson 2007 ⁴⁷	Sweden	IFG-WHO (14%)	5047 (66%)	57.5 (46-68)	10.7	Yes	CVD incidence	Yes
Pankow 2007 ⁴⁸	United States	IFG-ADA (42%) IGT (31%) IFG-ADA and IGT (17%)	6888 (47%)	62.3(52-75)	6.3	Yes	CHD incidence All-cause mortality	No
Rijkelijhuizen 2007 ¹⁰	Netherlands	IFG-ADA (33.2%) IFG-WHO (10.1%)	1428 (46%)	60.5 (50-75)	6.4	No	CVD mortality All-cause mortality	No*
Wang 2007 a ⁴⁹	Finland	IFG-ADA (NA) IFG-WHO (NA) IGT (NA)	1025 (63.2%)	NA (65-74)	13.5	No	CVD mortality	No
Wang 2007 ⁵⁰	China	IFG-ADA (56.5%) IFG-WHO (28.3%) IGT (24.6%)	541 (43.4%)	47.8 (≥25)	5	Yes	CHD incidence	No
Chien 2008 ⁵¹	China	IFG-WHO (NA) IGT (NA)	2165(56%)	54.2 (≥35)	10.5	Yes	CVD incidence	No
Jin 2008 ^{52 †}	China	IFG-ADA (11.4%)	1911 (0%)	72.5 (6.0)	9.5	No	CVD mortality All-cause mortality	Yes
Kim 2008 ⁵³	India	IFG-ADA (10%)	2993 (58%)	43 (≥35)	10.4	No	CHD mortality	No

Study	Country/ Region	Prediabetes definition and prevalence (%)	Sample size (% women)	Age (y), average (range or SD)	Follow-up (y)	Baseline CVD excluded	Events for analysis	Possibility of enrolling patients with diabetes
Levitzky 2008 ¹¹	United States	IGT (5.5%) IFG-ADA and IGT (7.7%) IFG-ADA (NA)	4058 (53.3%)	48.5 (10)	4	Yes	CVD incidence	Yes
Oizumi 2008 ⁵⁴	Japan	IFG-WHO (NA) IGT (12.3%) IFG-WHO (5.8%)	2938 (56.1%)	57.2 (11)	9.7	Yes	CHD incidence CVD incidence	No
Tsai 2008 ⁵⁵	China	IFG-WHO (6.9%)	35259 (34.2%)	50.9 (≥40)	15	Yes	CHD incidence Stroke incidence CVD mortality All-cause mortality	Yes
Wannamethee 2008 ⁵⁶	UK	IFG-WHO (NA)	5128 (0%)	NA (40-59)	20	Yes	CHD incidence	Yes
Hyvarinen 2009 ⁵⁷	Finland	IFG-WHO (17.3%)	18360 (54.4%)	55.2 (25-90)	12.9	Yes	Stroke incidence	No
Sung 2009 ⁵⁸	Sweden	IGT (21.9%)						
	Korea	IFG-ADA (13.4%) IFG-WHO (3.9%)	652901 (0%)	43 (30-64)	8.8	Yes	Stroke incidence	Yes
Doi 2010 ⁵⁹	Japan	IFG-ADA (NA) IFG-WHO (NA)	2421(57%)	57.6 (40-79)	14	Yes	CHD incidence Ischemic stroke	No
Khang 2010 ¹²	Korea	IGT (NA) IFG-ADA (32%)	9791 (55.2%)	43.2 (15)	5.8	Yes	CVD incidence CHD incidence Stroke incidence	Yes
Kokubo 2010 ⁶⁰	Japan	IFG-ADA (28%)	5321 (53%)	55 (30-79)	11.7	Yes	CVD incidence CHD incidence Stroke incidence	Yes
Magliano 2010 ⁶¹	Mauritius	IFG-WHO (5.7%) IGT (20.6%)	9559 (NA)	54.4 (20-82)	15.1	No	CVD mortality All-cause mortality	No
Skriver 2010 ⁶²	Denmark	HbA1c-NICE (4.1%)	25184 (53.0%)	54.9 (40-69)	6.8	No	All-cause mortality	No*
Sui 2011 ⁶³	United States	IFG-WHO (12.8%)	43,933(0)	44.3 (9.9)	17.7	Yes	Stroke incidence Stroke mortality	Yes
Yeboah 2011 ⁶⁴	United States	IFG-ADA (13.9%)	6753 (52.9%)	62.2 (45-84)	7.5	Yes	CVD incidence	Yes

Study	Country/ Region	Prediabetes definition and prevalence (%)	Sample size (% women)	Age (y), average (range or SD)	Follow-up (y)	Baseline CVD excluded	Events for analysis	Possibility of enrolling patients with diabetes
Ma 2012 ⁶⁵	Korea	IFG-ADA (15.9%)	16048 (60%)	55.7 (≥20)	9.4	No	CHD incidence Stroke incidence All-cause mortality CVD mortality	Yes
Deedwania 2013 ⁶⁶	United States	IFG-ADA (47%)	4602(57%)	73 (≥65)	13	No	CHD mortality Stroke mortality CVD mortality	No*
Kim 2013 ⁶⁷	Korea	IFG-ADA (17.7%)	408,022 (39.3%)	45.5 (≥20)	9.4	Yes	All-cause mortality CHD incidence Stroke incidence CVD incidence	Yes
Laukkanen 2013 ⁹	Finland	IFG-ADA (20.1%)	2641 (0%)	52.9 (40-62)	18.8	No	CHD incidence Stroke incidence CVD mortality	Yes
Onat 2013 ⁶⁸	Turky	IFG-WHO (5.9%) IGT (4.7%)	2619 (51.3%)	47.8 (11.8)	7.2	Yes	All-cause mortality CHD incidence	No
Schöttker 2013 ⁶⁹	German	IFG-ADA (21.6%) HbA1c-ADA (37.3%)	8365 (55.1%)	62 (50-74)	7.9	Yes	CVD incidence	Yes
Selvin 2014 ⁷⁰	United States	IFG-ADA (48.7%) HbA1c-ADA (23.1%)	9051 (58.4%)	56.6 (5.6)	6	Yes	CVD incidence All-cause mortality	Yes
Eastwood 2015 ⁷¹	South Asia Europe	HbA1c-ADA (31.9%) HbA1c-NICE (10.7%)	2475 (16.2%)	52 (40-69)	20	Yes	CVD incidence CHD incidence Stroke incidence	No
Gordon-Dseagu 2015 ⁷²	UK	HbA1c-ADA (24%)	22106 (54%)	52 (17.7)	7	No	All-cause mortality CVD mortality	Yes
Paprott 2015 ⁷³	German	HbA1c-ADA (27.8%)	6,299 (49%)	45.5 (18-79)	11.6	Yes	All-cause mortality	Yes

Study	Country/ Region	Prediabetes definition and prevalence (%)	Sample size (% women)	Age (y), average (range or SD)	Follow-up (y)	Baseline CVD excluded	Events for analysis	Possibility of enrolling patients with diabetes
Samaras 2015 ⁷⁴	Australia	IFG-ADA (40.7%) IFG-WHO (NA)	945 (54%)	78.6 (4.7)	2	No	All-cause mortality CHD incidence Stroke incidence	Yes
Kim 2016 ⁷⁵	Korea	IFG-ADA (26.4%) HbA1c-ADA (19.3%)	76434 (42.8%)	47.5 (10.5)	3.1	Yes	CVD incidence CHD incidence Stroke incidence All-cause mortality	Yes

* OGTT was performed in this study to exclude diabetes; † article in Chinese

CHD=coronary heart disease; CVD=cardiovascular disease; DECODE=Diabetes Epidemiology: Collaborative analysis Of Diagnostic criteria in Europe; HDL-C=high-density lipoprotein cholesterol; HbA1c-ADA=prediabetes defined as raised HbA1c according to the American Diabetes Association criteria (39-47 mmol/mol); HbA1c-NICE=prediabetes defined as raised HbA1c according to the NICE guidance (42-47 mmol/mol); IFG-ADA=impaired fasting glucose according to the American Diabetes Association criteria (fasting plasma glucose of 5.6-6.9 mmol/L); IFG-WHO=impaired fasting glucose according to the World Health Organization criteria (fasting plasma glucose of 6.1-6.9 mmol/L); IGT=impaired glucose tolerance; NA=not available; SD=standard deviation

Table B. Quality Assessment of the Included Studies

Study	Selection (stars awarded)	(stars	Comparability (stars awarded)	Outcome (stars awarded)	Quality (total stars awarded)
Stengard 1992 ²⁸	3		2	3	Good (8)
Barzilay 1999 ²⁹	4		2	1	Good (7)
DECODE 2001 ³⁰	3		2	3	Good (8)
Mazza 2001 ³²	3		1	2	Fair (6)
Saydah 2001 ³²	3		2	3	Good (8)
Henry 2002 ³³	4		2	3	Good (9)
Rodriguez 2002 ³⁴	4		2	2	Good (8)
Lu 2003 ³⁵	3		2	3	Good (8)
Hunt 2004 ³⁶	4		1	3	Good (8)
Nakagami 2004 ³⁷	3		2	2	Good (7)
Nakanishi 2004 ³⁸	4		2	2	Good (8)
Tai 2004 ³⁹	4		1	2	Good (7)
Hiltunen 2005 ⁴⁰	3		2	2	Good (7)
McNeill 2005 ⁴¹	3		2	2	Good (7)
Wild 2005 ⁴²	3		1	3	Good (7)
Kaarisalo 2006 ⁴³	4		1	2	Good (7)
McNeill 2006 ⁴⁴	4		1	3	Good (8)
Palmieri 2006 ⁴⁵	3		2	2	Good (7)
Barr 2007 ⁴⁶	4		1	2	Good (7)
Liu 2007 ⁸	4		1	3	Good (8)
Nilsson 2007 ⁴⁷	4		2	2	Good (8)
Pankow 2007 ⁴⁸	3		1	3	Good (7)
Rijkelijhuizen 2007 ¹⁰	3		1	2	Fair (6)
Wang 2007 a ⁴⁹	4		1	3	Good (8)
Wang 2007 ⁵⁰	4		1	2	Good (7)
Chien 2008 ⁵¹	3		2	2	Good (7)
Jin 2008 ⁵²	3		1	2	Fair (6)

Study	Selection awarded)	(stars	Comparability (stars awarded)	Outcome (stars awarded)	Quality (total stars awarded)
Kim 2008 ⁵³	4		2	2	Good (8)
Levitzky 2008 ¹¹	4		1	2	Good (7)
Oizumi 2008 ⁵⁴	2		1	2	Fair (5)
Tsai 2008 ⁵⁵	2		1	2	Fair (5)
Wannamethee 2008 ⁵⁶	4		1	2	Good (7)
Hyvarinen 2009 ⁵⁷	4		2	2	Good (8)
Sung 2009 ⁵⁸	4		2	2	Good (8)
Doi 2010 ⁵⁹	4		2	2	Good (8)
Khang 2010 ¹²	4		2	2	Good (8)
Kokubo 2010 ⁶⁰	4		2	3	Good (9)
Magliano 2010 ⁶¹	3		1	2	Fair (6)
Skriver 2010 ⁶²	3		2	2	Good (7)
Sui 2011 ⁶³	4		2	2	Good (8)
Yeboah 2011 ⁶⁴	4		2	3	Good (9)
Ma 2012 ⁶⁵	3		1	2	Fair (6)
Deedwania 2013 ⁶⁶	4		2	2	Good (8)
Kim 2013 ⁶⁷	4		2	2	Good (8)
Laukkanen 2013 ⁹	3		2	2	Good (7)
Onat 2013 ⁶⁸	4		2	2	Good (8)
Schöttker 2013 ⁶⁹	4		2	3	Good (9)
Selvin 2014 ⁷⁰	4		2	3	Good (9)
Eastwood 2015 ⁷¹	4		2	2	Good (8)
Gordon-Dseagu 2015 ⁷²	3		1	2	Fair (6)
Paprott 2015 ⁷³	4		2	2	Good (8)
Samaras 2015 ⁷⁴	3		2	1	Fair (6)
Kim 2016 ⁷⁵	4		2	1	Good (7)

Table C. Confounders Adjusted and Quality Assessment of the Included Studies

Study	Confounder adjusted	Adequate adjustment†
Stengard 1992 ²⁸	Age, BMI, hypertension, smoking, TC, HDL-C and functional capacity	Yes
Barzilay 1999 ²⁹	Age, sex, ethnic group, smoking, BMI, LDL-C and hypertension.	Yes
DECODE 2001 ³⁰	Age, sex, center, TC, BMI, SBP, smoking	Yes
Mazza 2001 ³²	Age, historical stroke and CHD, hypertension, SBP, pulse pressure, atrial fibrillation, LV hypertrophy, uric acid, smoking and serum potassium and sodium	No
Saydah 2001 ³²	Age, sex, race, education, smoking, physical activity , BMI, SBP, TC:HDL-C ratio	Yes
Henry 2002 ³³	Age, TC, TG, BMI, smoking and BP	Yes
Rodriguez 2002 ³⁴	Age, BMI, waist-to-hip ratio, physical activity, hypertension, TG, HDL-C, fibrinogen	Yes
Lu 2003 ³⁵	Age, sex, BMI, physical activity, study center, smoking, alcohol consumption, hypertension, insulin, TC, HDL-C, and TG.	Yes
Hunt 2004 ³⁶	Age, gender, and ethnic group	No
Nakagami 2004 ³⁷	Age, sex, cohort, BMI, SBP, TC and smoking.	Yes
Nakanishi 2004 ³⁸	Age, family history of diabetes, alcohol consumption, cigarette smoking, and all other components of the metabolic syndrome	Yes
Tai 2004 ³⁹	Age, sex, ethnicity	No
Hiltunen 2005 ⁴⁰	Age, gender, presence of CVD, hypertension, BMI, smoking, physical activity and self-perceived health	Yes
McNeill 2005 ⁴¹	Age, sex, hypertension, TC:HDL-C ratio, TG, smoking, baseline CVD.	Yes
Wild 2005 ⁴²	Sex, previous stroke/TIA, perceived health status, hypertension, BP, angina pectoris, previous MI, cardiac failure, atrial fibrillation, claudication, acetylsalicylic acid in use, smoking	No
Kaarisalo 2006 ⁴³	Age, sex, race	No
McNeill 2006 ⁴⁴	Age, sex, center	No
Palmieri 2006 ⁴⁵	Age, sex, previously CVD, smoking, hypertension, waist circumference, lipid-lowering medication use, and TC:HDL-C ratio.	Yes
Barr 2007 ⁴⁶	Age, sex, smoking, CVD family history, and TC	No
Liu 2007 ⁸	Age, sex	No
Nilsson 2007 ⁴⁷	Age, sex, race, center, smoking, hypertension, LDL-C, HDL-C, TG, lipid-lowering medications, BMI, and waist circumference.	Yes

Study	Confounder adjusted	Adequate adjustment†
Pankow 2007 ⁴⁸	Age, sex	No
Rijkelloshuizen 2007 ¹⁰	Age, sex, history of MI and stroke, current smoking, consumption of alcohol, physical activity and TC.	No
Wang 2007 a ⁴⁹	Age, sex, education, occupation, smoking, diabetic family history, and TC	No
Wang 2007 ⁵⁰	Age, sex, BMI, smoking, alcohol, marital status, education, occupation, exercise, family history of CHD	No
Chien 2008 ⁵¹	Age, post load plasma glucose, BMI, SBP, DBP, HDL-C, TC, TG	Yes
Jin 2008 ⁵²	Age, sex	No
Kim 2008 ⁵³	Age, sex, SBP, hypertension treatment, TC/HDL ratio, smoking, and BMI	Yes
Levitzky 2008 ¹¹	Age, sex, hypertension	No
Oizumi 2008 ⁵⁴	Age, gender, and smoking status	No
Tsai 2008 ⁵⁵	Age, gender, and smoking status	No
Wannamethee 2008 ⁵⁶	Age, smoking, social class, physical activity and alcohol intake	No
Hyvarinen 2009 ⁵⁷	Age, sex, center, mean arterial pressure, BMI, TC, smoking	Yes
Sung 2009 ⁵⁸	Age, height, smoking, alcohol consumption, regular exercise, level of monthly salary, area of residence, BP, TC, and BMI.	Yes
Doi 2010 ⁵⁹	Age, sex, SBP, electrocardiogram abnormalities, BMI, TC, HDL-C, smoking, alcohol intake, and regular exercise.	Yes
Khang 2010 ¹²	Age, sex, central obesity, hypertriglyceridemia, HDL-C, hypertension	Yes
Kokubo 2010 ⁶⁰	Age, sex, BMI, hypertension, hyperlipidemia, and smoking and drinking status	Yes
Magliano 2010 ⁶¹	Prior CVD, education, sex, hypertension, waist and hip circumference, smoking, HDL-C, TG, and TC.	No
Skriver 2010 ⁶²	Age, sex, BMI, smoking, SBP, history of ischaemic heart disease, cerebrovascular disease and cancer	Yes
Sui 2011 ⁶³	Age, year of examination, smoking, alcohol intake, family history of CVD, and survey indicator, BMI, TC, abnormal ECG, and hypertension.	Yes
Yeboah 2011 ⁶⁴	Age, sex, race/ethnicity, BMI, SBP, TC, HDL-C, TG, smoking, BP medications and statin use.	Yes
Ma 2012 ⁶⁵	Age, sex, educational, alcohol consumption, history of hypertension and BMI	No
Deedwania 2013 ⁶⁶	Age, sex, race/ethnicity, married, education, income, BMI, activities of daily living, smoking, alcohol use, ankle arm index ratio, hemoglobin, TC, albumin, uric acid, C-reactive protein, serum insulin, LV hypertrophy, atrial fibrillation, bundle branch block, LV systolic dysfunction, chronic	Yes

Study	Confounder adjusted	Adequate adjustment†
Kim 2013 ⁶⁷	disease and medicine Age, SBP, antihypertensive medication, LDL-C, HDL-C, current smoking, BMI, and family history of CVD.	Yes
Laukkanen 2013 ⁹	Age, BMI, SBP, LDL-C, smoking, alcohol consumption, prevalent CHD, and family history of CHD	Yes
Onat 2013 ⁶⁸	Age, sex, SBP, non-HDL-C, waist circumference, smoking, and C-reactive protein	Yes
Schöttker 2013 ⁶⁹	Age, sex, SBP, current smoking and TC/HDL-C ratio	Yes
Selvin 2014 ⁷⁰	Age, sex, race-center, BMI, C-reactive protein, smoking, SBP, LDL-C, HDL-C, TG, estimated glomerular filtration rate, current use of hypertension medication, current lipid-lowering medication use, alcohol use and left ventricular hypertrophy.	Yes
Eastwood 2015 ⁷¹	Age, sex, smoking, total-to-HDL cholesterol ratio, waist-to-hip ratio, SBP, antihypertensive treatment.	Yes
Gordon-Dseagu 2015 ⁷²	Age, sex, smoking, BMI, socioeconomic status	No
Paprott 2015 ⁷³	Age, sex, educational level, smoking, sport activity, alcohol consumption, BMI, waist circumference, history of myocardial infarction, stroke, or cancer, and history of hypertension or hyperlipidemia	Yes
Samaras 2015 ⁷⁴	Age, sex, hypertension, BMI, smoking	Yes
Kim 2016 ⁷⁵	Age, sex, hypertension, LDL-C, HDL-C, smoking, family history of CVD, and BMI	Yes

†Adequate adjustment denoted adjustment of at least five of six confounders including sex, age, hypertension or blood pressure or antihypertensive treatment, body mass index or other measure of overweight/obesity, cholesterol, and smoking

BMI: body mass index; BP: blood pressure; HDL-C: high-density lipoprotein cholesterol; LDL-C: low-density lipoprotein cholesterol; LV: left ventricular; CHD: coronary heart disease; CVD: cardiovascular disease; MI: myocardial infarction; SBP: systolic blood pressure; TC: total cholesterol; TG: triglycerides

Table D. Definition of composite cardiovascular disease in the Included Studies

Study	Definition of composite cardiovascular disease
Stengard 1992 ²⁸	CHD and stroke
Barzilay 1999 ²⁹	CHD and stroke
DECODE 2001 ³⁰	ICD-9 codes: 401-448
Saydah 2001 ³²	ICD-9 codes: 401-448
Henry 2002 ³³	ICD-9 codes: 390-459
Rodriguez 2002 ³⁴	CHD, stroke and sudden death
Lu 2003 ³⁵	ICD-9 codes: 390-448
Hunt 2004 ³⁶	ICD-9 codes: 401-405, 410-414, 420-429 (with the exception of 427.5), 430-439, or 440-447 .
Nakagami 2004 ³⁷	ICD-9 codes: 401-448
Nakanishi 2004 ³⁸	CHD and stroke
Tai 2004 ³⁹	ICD-9 codes: 401-448
Wild 2005 ⁴²	ICD-9 codes: 410-414, 430-438, 440-445 and ICD-10 codes: I21-25, I60-73
Barr 2007 ⁴⁶	ICD-10 codes: I10-I25, I46.1, I48, I50-I99, or R96
Liu 2007 ⁸	CHD, stroke and sudden death
Nilsson 2007 ⁴⁷	ICD-9 codes: 410-414, 430, 431, 434, 436
Rijkelijkhuisen 2007 ¹⁰	ICD-9 codes: 390-459, 798
Wang 2007 a ⁴⁹	ICD-9 codes: 390-459
Chien 2008 ⁵¹	CHD and stroke
Jin 2008 ^{52 †}	CHD death, sudden death, stroke death,
Levitzky 2008 ¹¹	CHD, stroke, transient ischemic attack (TIA), intermittent claudication, congestive heart failure, or CVD death
Oizumi 2008 ⁵⁴	CHD and stroke
Tsai 2008 ⁵⁵	ICD-9 codes: 390-459
Khang 2010 ¹²	ICD-10 codes: I20-I25, I60-I69
Kokubo 2010 ⁶⁰	CHD and stroke
Magliano 2010 ⁶¹	Deaths categorized as cardiac, cerebrovascular, hypertension, or renal failure the primary cause
Yeboah 2011 ⁶⁴	MI, definite angina, probable angina (if followed by coronary artery bypass grafting and percutaneous coronary intervention), resuscitated cardiac arrest, stroke, stroke death, CHD death or other CVD death
Ma 2012 ⁶⁵	ICD-10 codes: I00-I99
Deedwania 2013 ⁶⁶	MI, angina pectoris, stroke, and peripheral artery disease.
Kim 2013 ⁶⁷	ICD-10 codes: I10-I15, I20-I25, I44-I51, I60-69, I70-I74, R96),
Laukkanen 2013 ⁹	ICD-9 codes: 798
Selvin 2014 ⁶⁸	CHD and stroke
Schöttker 2013 ⁶⁹	MI, stroke or cardiovascular death (ICD-10 codes: I00-I99).
Eastwood 2015 ⁷¹	CHD and stroke
Gordon-Dseagu 2015 ⁷²	ICD-9 codes:390–459; ICD-10 codes:I00-I99
Kim 2016 ⁷⁵	Cardiovascular death (ICD-10 codes: I00-99 and R96), MI, stroke, coronary revascularization, hospitalization or outpatient visit with ischemic heart disease, other vascular disease, or congestive heart failure.

CHD: coronary heart disease; CVD: cardiovascular disease; ICD: International Classification of Diseases; MI: myocardial infarction

Table E. Subgroup Analyses of the Association between Prediabetes and All-Cause Mortality

Subgroups	IFG-ADA			IFG-WHO			IGT		
	No of studies	RR (95% CI)	P *	No of studies	RR (95% CI)	P *	No of studies	RR (95% CI)	P *
Ethnicity			0.36			0.79			0.89
Asians	2	1.24 (1.01, 1.53)		3	1.14 (1.02, 1.27)		3	1.30 (1.08, 1.56)	
Non-Asians	7	1.11 (0.98, 1.25)		10	1.16 (1.03, 1.31)		8	1.32 (1.23, 1.41)	
Sex			-			0.68			0.63
Male	2	1.28 (1.11, 1.46)		4	1.14 (1.03, 1.26)		5	1.34 (1.21, 1.48)	
Female	0	-		2	1.07 (0.79, 1.44)		4	1.28 (1.11, 1.48)	
Participant's average age			0.009			0.88			0.07
<55 years	3	1.28 (1.13, 1.46)		7	1.12 (1.05, 1.21)		5	1.36 (1.26, 1.47)	
≥55 years	6	1.04 (0.96, 1.13)		5	1.15 (0.91, 1.45)		5	1.19 (1.04, 1.35)	
Follow-up duration			0.86			0.38			0.47
<10 years	7	1.12 (0.99, 1.27)		8	1.21 (1.04, 1.41)		7	1.34 (1.24, 1.44)	
≥10 years	2	1.15 (0.91, 1.46)		5	1.12 (1.01, 1.23)		4	1.27 (1.13, 1.42)	
Possibility of enrolling patients with diabetes			0.35			0.65			0.64
None	3	1.08 (0.89, 1.31)		9	1.13 (1.02, 1.25)		10	1.32 (1.23, 1.41)	
Might enrolled	6	1.20 (1.08, 1.34)		4	1.17 (1.05, 1.31)		1	1.17 (0.71, 1.93)	
Exclusion of individuals with baseline CVD			0.14			0.73			0.87
Yes	4	1.02 (0.86, 1.19)		4	1.17 (1.05, 1.30)		2	1.34 (1.0, 1.81)	
No	5	1.19 (1.03, 1.38)		9	1.13 (1.02, 1.27)		9	1.31 (1.23, 1.4)	
Adjustment of confounders			0.11			0.81			0.91
Adequate [†]	8	1.09 (1.02, 1.18)		9	1.14 (1.04, 1.25)		10	1.32 (1.23, 1.41)	
Not adequate	1	1.41 (1.04, 1.91)		4	1.17 (0.98, 1.39)		1	1.30 (1.11, 1.53)	

IFG-ADA=impaired fasting glucose according to the American Diabetes Association definition (fasting glucose 5.6-6.9 mmol/L); IFG-WHO=impaired fasting glucose according to the World Health Organization definition (fasting glucose 6.1-6.9 mmol/L); IGT=impaired glucose tolerance

* For heterogeneity among subgroups.

† Adequate adjustment denoted adjustment of at least five of six confounders including sex, age, hypertension or blood pressure or antihypertensive treatment, body mass index or other measure of overweight/obesity, cholesterol, and smoking

Table F: Subgroup Analyses of the Association between Prediabetes and Risk of Composite Cardiovascular Diseases

Subgroups	IFG-ADA			IFG-WHO			IGT		
	No of studies	RR (95% CI)	P*	No of studies	RR (95% CI)	P*	No of studies	RR (95% CI)	P*
Ethnicity			0.65			0.92			0.74
Asians	7	1.11 (1.01, 1.26)		6	1.24 (0.97, 1.58)		5	1.33 (1.12, 1.58)	
Non-Asians	8	1.15 (1.04, 1.27)		12	1.26 (1.10, 1.43)		8	1.29 (1.17, 1.42)	
Sex			0.40			0.91			0.66
Male	8	1.09 (1.01, 1.18)		6	1.17 (1.03, 1.33)		4	1.34 (1.17, 1.53)	
Female	7	1.18 (1.00, 1.39)		3	1.19 (0.86, 1.65)		3	1.27 (1.04, 1.55)	
Participant's average age			0.49			0.78			0.98
<55 years	6	1.11 (1.01, 1.21)		9	1.27 (1.04, 1.55)		6	1.29 (1.16, 1.44)	
≥55 years	9	1.16 (1.04, 1.30)		8	1.22 (1.04, 1.44)		6	1.29 (1.13, 1.46)	
Follow-up duration			0.15			0.54			0.87
<10 years	10	1.08 (1.01, 1.15)		10	1.22 (1.02, 1.44)		8	1.31 (1.18, 1.44)	
≥10 years	5	1.22 (1.05, 1.41)		8	1.31 (1.12, 1.52)		5	1.29 (1.12, 1.48)	
Possibility of enrolling patients with diabetes			0.19			0.31			0.65
None	2	1.31 (1.04, 1.65)		12	1.22 (1.05, 1.42)		12	1.30 (1.20, 1.42)	
Might enrolled	12	1.12 (1.04, 1.20)		6	1.36 (1.08, 1.57)		1	1.13 (0.62, 2.06)	
Exclusion of individuals with baseline CVD			0.22			0.1			0.74
Yes	9	1.10 (1.03, 1.18)		9	1.38 (1.23, 1.55)		5	1.27 (1.09, 1.49)	
No	6	1.27 (1.02, 1.58)		9	1.15 (0.97, 1.38)		8	1.31 (1.19, 1.45)	
CVD endpoint			0.22			0.22			0.90
CVD incidence	9	1.10 (1.03, 1.18)		5	1.39 (1.16, 1.68)		4	1.29 (1.11, 1.50)	
CVD mortality	6	1.27 (1.02, 1.58)		13	1.20 (1.05, 1.38)		9	1.30 (1.18, 1.44)	
Adjustment of confounders			0.05			0.13			0.34
Adequate [†]	11	1.09 (1.02, 1.18)		10	1.16 (0.98, 1.37)		8	1.26 (1.14, 1.40)	
Not adequate	4	1.26 (1.12, 1.43)		8	1.37 (1.19, 1.58)		4	1.38 (1.18, 1.61)	

CVD=cardiovascular disease; IFG-ADA=impaired fasting glucose according to the American Diabetes Association definition (fasting glucose 5.6-6.9 mmol/L); IFG WHO= impaired fasting glucose according to the World Health Organization definition (fasting glucose 6.1-6.9 mmol/L); IGT=impaired glucose tolerance

* For heterogeneity among subgroups.

† Adequate adjustment denoted adjustment of at least five of six confounders including sex, age, hypertension or blood pressure or antihypertensive treatment, body mass index or other measure of overweight/obesity, cholesterol, and smoking