


A systematic review and meta-analysis of retrograde type A aortic dissection after thoracic endovascular aortic repair in patients with type B aortic dissection

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Abstract

Background: Retrograde type A dissection (RTAD) is a devastating complication of thoracic endovascular repair (TEVAR) with low incidence but high mortality. The objective of this study is to report the incidence, mortality, potential risk factors, clinical manifestation and diagnostic modalities, and medical and surgical treatments.

Methods: A systematic review and single-arm and two-arm meta-analyses evaluated all published reports of RTAD post-TEVAR through January 2021. All study types were included, except study protocols and animal studies, without time restrictions. Outcomes of interest were procedural data (implanted stent-grafts type, and proximal stent-graft oversizing), the incidence of RTAD, associated mortality rate, clinical manifestations, diagnostic workups and therapeutic management.

Results: RTAD occurred in 285 out of 10,600 patients: an estimated RTAD incidence of 2.3% (95% CI: 1.9–2.8); incidence of early RTAD was approximately 1.8 times higher than late. Wilcoxon signed-rank testing showed that the proportion of RTAD patients with acute type B aortic dissection (TBAD) was significantly higher than those with chronic TBAD ($P = .008$). Pooled meta-analysis showed that the incidence of RTAD with proximal bare stent TEVAR was 2.1-fold higher than with non-bare stents: risk ratio was 1.55 (95% CI: 0.87–2.75; $P = .13$). Single arm meta-analysis estimated a mortality rate of 42.2% (95% CI: 32.5–51.8), with an I^2 heterogeneity of 70.11% ($P < .001$).

Conclusion: RTAD is rare after TEVAR but with high mortality, especially in the first month post-TEVAR with acute TBAD patients at greater risk as well as those treated with proximal bare stent endografts.

Abbreviations: CI = confidence interval, CT = computed tomography, ICU = intensive care unit, RR = risk ratio, RTAD = retrograde type A dissection, TBAD = type B aortic dissection, TEVAR = thoracic endovascular repair.

Keywords: complication, meta-analysis, retrograde type A aortic dissection, TEVAR.

1. Introduction

Aortic dissection generally has a high rate of mortality if untreated; with Type A aortic dissection particularly, 30-day mortality can be as high as 90%.^[1] The true incidence of aortic

dissection is not well known, but with the advent of new diagnostic modalities over the last decade, estimations have dramatically risen.^[2,3] Annually, 5 to 10 people per million experience an aortic dissection in the United States with 43,000 to 47,000

SA-H-A-S and NH contributed equally to this work.

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The datasets generated during and/or analyzed during the current study are available from the corresponding author on reasonable request.

Since our study is based on already published literature with no interaction with human subjects, no issues related to medical ethics were needed to be reported.

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lives claimed due to the involvement of the aorta and its branches.^[4,5]

The condition is conventionally classified as Stanford Types A or B, with the latter involving the descending aorta. New classifications – such as TEM (Type of dissection, location the primary Entry, and Malperfusion) and the Society for Vascular Surgeons reporting standards – have further clarified the extent of the disease process and improved awareness of the disease mechanism to guide decision making and predict outcomes.^[6,7] Acute Type B aortic dissection (TBAD) is an uncommon condition involving the descending aorta that remains a challenging problem for cardiothoracic and vascular surgeons as well as interventional radiologists whereas treatment of chronic TBAD can vary between medical and surgical therapies.^[8–10]

Conventionally, patients with uncomplicated TBAD receive medical treatment, while evidence progressively support thoracic endovascular aortic repair (TEVAR) as the preferred treatment for complicated and some high-risk TBAD according to Society for Vascular Surgeons guidelines.^[7,11] While endovascular techniques were initially used for patients not indicated for conventional surgery, indications have rapidly expanded owing to recent clinical progress over the last decades.^[12] Increasing evidence shows positive TEVAR outcomes with acceptable protection against aorta-related death in mid-term follow-up. TEVAR stabilizes the dissected aorta and prevents late complications by expanding the true lumen, inducing both false lumen thrombosis and aortic wall remodeling. In comparison with traditional open aortic surgery, TEVAR has the benefits of fewer complications, smaller incisions, and shorter length of hospital stay which explains the reason that it is currently the preferred treatment for complicated TBAD.^[13]

TEVAR is still linked with major complications such as acute or delayed retrograde Type A dissection (RTAD), stroke, bowel infarction, access-related complications, paraplegia, endoleaks, limb ischemia, or wound infection.^[14] RTAD is a devastating complication of this procedure with a low incidence, but mortality rates exceed 40%.^[15] A wide range of studies on RTAD post-TEVAR have reported small numbers of patients with unclear diagnostic and therapeutic approaches. Different etiologies have been proposed for RTAD but is essentially due to unfavorable interaction between the stent-graft and dissecting membrane that can produce a new primary entry tear and lead to rupture of the membrane. Interpretation is complicated by heterogeneity of data quality, definitions and the reported parameters; as well by its broader relation to any stent graft-induced aortic wall injury and to other iatrogenic injury in non-dissections.^[13,16–18] RTAD is also sometimes referred to as proximal SINE (to complement distal stent-graft-induced new entry).^[19]

We conducted this comprehensive systematic review and single-arm and two-arm meta-analyses to identify all published reports on RTAD post-TEVAR with the intention of recording the incidence, mortality, potential risk factors, clinical manifestation and diagnostic modalities, and medical and surgical treatments. The findings might assist in designing appropriate clinical strategies to minimize occurrence and diagnose and treat this complication early and effectively in the hope of improving future procedural safety and outcomes.

2. Methods

This is a systematic review carried out according to Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guidelines (see Table S1, Supplemental Digital Content, <http://links.lww.com/MD/I478>, which illustrates PRISMA 2020 checklist).^[20] We used the PICOS strategy (population, intervention, comparison, outcomes, and design of studies) to formulate the research question and eligibility criteria:

Population: patients with Type B aortic dissection

Intervention: thoracic endovascular aortic repair

Comparators: none

Outcome: incidence of RTAD, re-intervention and its types, mortality of RTAD

Study design: all study designs except for study protocols, animal studies.

To eliminate the risk of analyzing the same patients more than once, the studies were assessed and duplicate publications and overlapping reports were removed. Extensive effort was made to minimize the impact of covert duplicate or metachronous re-publication from the same groups or patient cohorts; for these cases, only the latest report was included.

The search was conducted in PubMed, Cochrane Central, Embase, and Web of Science databases through January 2021.^[21,22] The search terms included “TEVAR,” “retrograde dissection,” “thoracic stent-graft,” “endograft,” and “graft” with the Boolean operator “OR,” was restricted to English-language results and with no limits on date of publication. All retrieved results were assessed and screened to obtain additional relevant articles not indexed in common databases.

To be included in the meta-analysis, publications had to meet all the following inclusion criteria: (1) Articles reporting complications of RTAD post-TEVAR among those who underwent endovascular repair or hybrid repair of thoracic aortic pathology; (2) Diagnosis of aortic pathology made by computed tomography (CT) imaging of the thorax, abdomen, or pelvis; (3) Series with more than 5 patients with TEVAR; (4) Demographic data and comorbidities of the patients; (5) At least one of the basic outcome criteria (number of patients with TEVAR, number of patients with RTAD, or mortality of RTAD).

After first screening of titles and abstracts in selected electronic databases, the full texts of appropriate studies were evaluated and their data were extracted by three investigators (SAHS, NH, and MMO) independently. Discrepancies among these investigators were resolved through discussions with a senior author (HE). The following data for each study were extracted: study characteristics, patient characteristics, studies quality, aortic pathology, procedural data (implanted stent-grafts type, and proximal stent-graft oversizing), mean follow-up period, number of patients with RTAD, re-intervention and its types, and RTAD mortality rate.

Since our study is based on already published literature with no interaction with human subjects, no issues related to medical ethics were needed to be reported.

2.1. Definition of extracted data

Regular and irregular imaging follow-up period was considered as ≥ 3 thoracic CTs after TEVAR and < 3 , respectively. Aortic dissection was described as an acute event if it occurred within the first 14 days from the onset of symptoms, and chronic beyond 14 days. Postoperative mortality was defined as all death events occurred during follow-up. Early RTAD or early mortality was considered if occurred within the first 3 months from the TEVAR procedure, while late RTAD or late mortality occurred after 3 months from the TEVAR procedure.^[23,24]

2.2. Statistical analysis

For the single-arm meta-analysis, analyses of proportions were conducted for data using a random effects model to calculate pooled incidences of RTAD and mortality rates and their confidence intervals (CI) using per protocol and intention to treat data when available. For the two-arm meta-analysis, dichotomous data were presented as risk ratios (RR) and continuous data as weighted mean differences. Summary effect measures were presented along with their corresponding 95% CIs. Statistical heterogeneity was evaluated with the I^2 statistic. P value between 0% and 25% indicates insignificant heterogeneity, 26% and 50% low heterogeneity, 51% and 75% moderate

heterogeneity, and 76% and 100% high heterogeneity. When I^2 was < 50%, a fixed-effects model was used and when it was > 50%, a random-effects model. For the analysis of other data that were not included in the meta-analysis, the data were analyzed using the statistical package IBM SPSS version 26.0 (Statistical Package for the Social Sciences, Chicago, IL). The categorical variables are expressed as proportions and frequencies. The continuous variables are summarized as mean ± standard-deviations. Also, in order to explore the independent nature of some categorical variables, Chi-square or exact Fisher test were used. Normality of numerical variables was checked using the Kolmogorov Smirnov test. *t*-test or Wilcoxon test were applied for comparing of two related groups. One-way ANOVA, Kruskal Wallis and Friedman tests also were implemented based on the normality test for more than two-group comparisons. A *P* value less than .05 was considered statistically significant in all analyses.

3. Results

The literature search yielded 1963 potentially eligible articles. After considering our selection criteria, 78 eligible clinical studies^[4,5,8,10,12,14,17,25-95] published between 2002 and 2020 were enrolled in the qualitative and quantitative analysis (Fig. 1). Of the total included records, 59, 10, and 9 studies were single-center, national multi-center, and international multi-center studies, respectively (Table 1). Most of the studies were conducted in Europe (31/78 studies; 39.7%) and Asia (26/78 studies; 33.3%). Sixteen studies (20.5%) were conducted in North America, one in South America, and four were

multi-continental studies. The studies were assigned into two categories according to the number of TBAD patients undergoing TEVAR during the study period. Thirty-nine studies (50%) with 1321 patients, and 39 (50%) with 9279 cases had < 50 and > 50 cases, respectively.

Table 2 summarizes the demographic and perioperative characteristics of 10,600 TBAD patients who underwent TEVAR. Patient populations ranged from 5 to 852, with a mean age of 57.4 years, 77.8% being male. Hypertension (83.4%) and smoking (47.7%) were the leading underlying diseases. Preoperative details are summarized in supporting information (see Table S2, Supplemental Digital Content, <http://links.lww.com/MD/I479>, which illustrates reported risk factors for RTAD). A majority of cases were acute TBAD: in 61/78 reports (*n* = 6741), 4049 cases (60%) were specified as acute TBAD; in 59 reports (*n* = 6686), 2997 cases (44.8%) were chronic TBAD. However, 17 and 19 studies, respectively, did not specify TBAD chronicity (see Table S3, Supplemental Digital Content, <http://links.lww.com/MD/I480>, which illustrates TBAD chronicity).

Of patients who experienced RTAD, mean age was 56.6 years and 85.7% were male. Hypertension (86.2%) was the most common comorbidity for RTAD, followed by smoking (65.6%), pulmonary disease (17.0%), Marfan syndrome (15.5%), renal impairment (14.6%), diabetes mellitus (14.2%), and coronary artery disease (12.5%) (Table 2).

Table 3 presents TEVAR details and the stent-grafts used in each study. From 50 enrolled studies, proximal bare stents were used in 3033 (66%) and proximal non-bare stents in 1569 cases (34%).

RTAD occurred in 285 cases out of 10,600 patients, representing an estimated RTAD incidence of 2.3% (95%

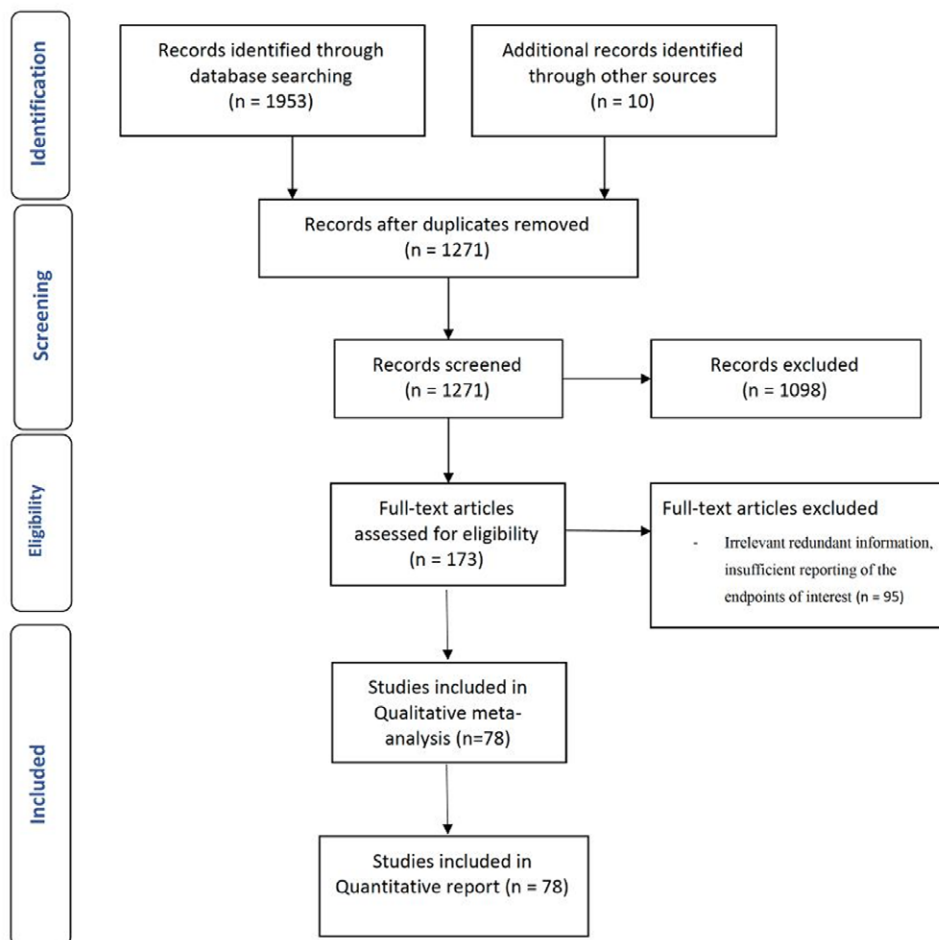


Figure 1. PRISMA flow chart of the study.

Table 1**Details and characteristics of studies reporting retrograde type A dissection after thoracic endovascular aortic repair.**

First author	Year	Duration	Geography	Center	Mean follow-up (mo)	TEVARs (n)	RTAD (n)	Age (yr)		Male sex (%)	
								TEVAR	RTAD	TEVAR	RTAD
Czermak ^[25]	2002	(1996–2001)	Austria	Innsbruck SC	17.8	5	1	51.12	43	NR	NR
Kato ^[26]	2002	(1997–2001)	Japan	Mie, Matsusaka SC	27	28	1	66.6	NR	22	NR
Palmer ^[27]	2002	(1999–2001)	Germany	Ulm SC	14	14	2	60.3	47.5	12	2
Fattori ^[28]	2003	(1997–2002)	Italy	Bologna SC	25	22	2	NR	NR	NR	NR
Grabenwoger ^[29]	2004	(1996–2003)	Austria	Vienna SC	NR	20	1	NR	NR	NR	NR
Hansen ^[30]	2004	(1998–2003)	USA	Torrance SC	24	24	1	69 (43–86)	NR	NR	NR
Lee ^[31]	2004	(1994–2002)	South Korea	Seoul SC	34	37	1	NR	NR	NR	NR
Dong Xu ^[32]	2005	(2001–2004)	China	Beijing SC	32	24	3	NR	NR	NR	NR
Fattori ^[33]	2006	(1996–2004)	Italy, Germany, France, Netherlands, etc.	Multicenter IMC	24	180	2	NR	NR	NR	NR
Duebener ^[34]	2007	(2000–2006)	Germany	Luebeck SC	38	13	1	59.5	NR	10	NR
Zipfel ^[35]	2007	(1999–2005)	Germany	Berlin SC	23	57	1	62	38	43	0
Kpodonu ^[36]	2008	(2000–2006)	USA	Pennsylvania NMC	NR	91	6	NR	69	NR	3
Neuhauser ^[14]	2008	(1997–2007)	Austria	Innsbruck SC	43	28	5	NR	65	NR	4
Dong ^[37]	2009	(2000–2007)	China	Shanghai SC	26	443	11	NR	43	NR	NR
Chiesa ^[38]	2011	(1999–2011)	Italy	Milan SC	NR	188	3	NR	NR	NR	NR
Kim ^[39]	2011	(2002–2009)	USA	Torrance SC	12.4	41	3	67.6	NR	31	NR
Oberhuber ^[40]	2011	(1999–2011)	Germany	Ulm SC	12.7	19	1	60	NR	17	NR
Parsa ^[41]	2011	(2005–2009)	USA	North Carolina SC	27	51	2	57	NR	37	NR
Wiedemann ^[4]	2013	(1996–2010)	Austria	Vienna SC	52	80	3	59	NR	58	NR
Lotfi ^[42]	2013	(1997–2011)	UK	London SC	15	11	3	NR	NR	NR	NR
Wiedemann ^[5]	2014	(1999–2011)	Austria, France, Italy, Spain, USA	Multicenter IMC	37	110	6	61	NR	86	NR
Faure ^[43]	2014	(2000–2011)	France	Montpellier SC	12.2	41	1	66	NR	34	NR
Idrees ^[44]	2014	(2000–2012)	USA	Cleveland, Ohio SC	48	766	15	NR	65	NR	NR
Zhang ^[45]	2014	(1998–2012)	China	Shanghai SC	58.4	252	2	54.1	NR	206	NR
Gorlitzer ^[46]	2012	(2005–2011)	Austria, Switzerland	Vienna, Bern IMC	NR	29	4	NR	62	NR	2
Huang ^[47]	2013	(2004–2011)	China	Guangzhou SC	NR	563	4	54.09	62.75	485	3
Cochemnec ^[48]	2013	(2004–2011)	France	Creteil SC	24.5	17	4	60	63.75	11	2
Shuyang Lu ^[49]	2012	(2006–2011)	China	Shanghai SC	34.79	419	9	NR	56.6	277	6
Yang ^[50]	2012	(2006–2011)	Taiwan	Taipei SC	24.1	61	1	62.7	NR	51	NR
Bunger ^[51]	2013	(2006–2012)	Germany	Rostock SC	27.9	45	1	59.9	55	38	1
Canaud ^[52]	2014	(2002–2012)	UK	London SC	NR	309	11	63.1	NR	248	NR
Lombardi ^[53]	2012	(2007–2012)	Italy, Germany, Australia, USA	Multicenter IMC	12	40	3	58	NR	28	NR
Jia ^[54]	2013	(2007–2010)	China	Beijing, Zhengzhou, Xinxiang NMC	28.5	208	3	52.1	NR	154	NR
Li ^[55]	2014	(2005–2012)	China	Beijing NMC	32.2	669	6	NR	41.2	NR	20
Hanna ^[56]	2014	(2005–2012)	USA	North Carolina SC	34.1	50	1	59	NR	36	NR
De Rango ^[57]	2014	(2005–2013)	Italy	Rome, Perugia NMC	29.2	104	4	69.8	NR	90	NR
Appoo ^[58]	2015	(2008–2012)	Canada	Alberta SC	72	16	0	63.8	NR	NR	NR
Desai ^[59]	2015	(2005–2012)	USA	Philadelphia SC	132	9	9	64.1	NR	80	NR
Kische ^[60]	2015	(2009–2015)	Germany	Berlin, Rostock NMC	25.6	35	1	63	NR	27	NR
Bockler ^[61]	2016	(2009–2010)	Germany, UK, Italy, Sweden	Multicenter IMC	24	24	1	NR	NR	NR	NR
Faure ^[62]	2016	(2005–2015)	France	Montpellier SC	24.3	33	1	65.1	62	26	1
Wang ^[63]	2016	(2005–2013)	China	Zhengzhou SC	32	360	5	52	51.8	304	4
Asaloumidis ^[64]	2017	(2000–2014)	Greece	Thessaloniki SC	74	40	2	65	NR	33	NR
Zhao Liu ^[65]	2017	(2008–2016)	China	Nanjing SC	30.5	58	6	57.3	NR	40	NR
Min-Hong Zhang ^[66]	2017	(2011–2013)	China	Beijing SC	26.4	85	3	64.3	NR	59	NR
Tjaden ^[67]	2018	(2010–2016)	USA, Europe, Brazil and Oceania	Multicenter IMC	26	264	6	62	NR	211	NR
Tao Ma ^[68]	2018	(2005–2013)	China, UK	Shanghai, London IMC	31.2	852	27	55	NR	720	NR
Laquian ^[69]	2018	(2011–2014)	USA	Florida, Alabama NMC	17.9	27	1	63	NR	17	NR
Chen ^[70]	2018	(2007–2014)	China	Hebei, Beijing NMC	17.9	167	1	NR	NR	112	NR
Piotr Buczkowski ^[71]	2019	(2007–2017)	Poland	Poznan SC	55	68	2	NR	NR	NR	NR
Eleshra ^[72]	2020	(2010–2017)	Germany	Hamburg SC	28	64	1	64.8	NR	49	NR
Fukushima ^[73]	2019	(2011–2017)	Japan	Chiba SC	14.2	24	0	67.7	NR	21	NR
Wang ^[74]	2019	(2013–2014)	USA	Multicenter IMC	1	397	6	60.4	NR	286	NR
Yammine ^[17]	2019	(2012–2017)	USA	North Carolina SC	14.25	186	15	61.6	61.5	112	8
Miura ^[75]	2019	(2013–2017)	Japan	Sapporo SC	19.6	22	0	63	NR	16	NR

(Continued)

Table 1
(Continued)

First author	Year	Duration	Geography	Center	Mean follow-up (mo)	TEVARs (n)	RTAD (n)	Age (yr)		Male sex (%)		
								TEVAR	RTAD	TEVAR	RTAD	
Chassin-Trubert ^[76]	2020	(2013–2019)	France	Montpellier	SC	26	17	0	NR	NR	NR	NR
Pellenc ^[77]	2019	(2015–2018)	France	Paris	SC	22	20	0	NR	NR	NR	NR
Jiechang Zhu ^[78]	2018	(2015–2016)	China	Tianjin	SC	6.95	20	0	53	NR	16	NR
Riesterer ^[79]	2018	(2002–2017)	Germany	Freiburg	SC	16	34	1	NR	NR	NR	NR
Giles ^[12]	2019	(2005–2016)	USA	Gainesville	SC	17	258	12	61.5	NR	203	NR
Kuo ^[80]	2019	(2006–2016)	USA	Los Angeles	SC	14	71	2	58.6	NR	52	NR
Joo ^[81]	2019	(1994–2017)	South Korea	Seoul	SC	NR	17	2	50.4	42	14	2
Cao ^[82]	2020	(2015–2018)	China	Beijing	SC	17.6	76	4	50.3	NR	51	NR
El-Beyrouti ^[83]	2020	(2018–2019)	Germany	Mainz, Tübingen	NMC	11.6	5	0	NR	NR	NR	NR
Charltonouw ^[84]	2018	(1999–2014)	USA	Houston	SC	51.6	43	3	NR	NR	NR	NR
Lou ^[85]	2020	(2012–2018)	USA	South Carolina	SC	36	91	3	52.6	NR	60	NR
Lee ^[86]	2020	(2003–2017)	South Korea	Seoul, Incheon and Cheonan	NMC	39.4	87	2	58.3	NR	62	NR
Oshi ^[87]	2020	(2009–2019)	Japan	Fukuoka	SC	39.2	40	1	66.5	NR	26	NR
Puech-Leao ^[88]	2020	(2004–2017)	Brazil	Sao Paulo	SC	57	42	4	59.1	NR	32	NR
Sobocinski ^[89]	2020	(2005–2015)	Sweden, France	Multicenter	IMC	1	41	2	58.8	NR	32	NR
Shuo Zhao ^[90]	2020	(2009–2018)	China	Shandong	SC	10.7	79	1	49.9	NR	61	NR
Bavaria ^[91]	2015	(2010–2012)	USA	Multicenter	NMC	12	50	2	57.2	NR	40	NR
Peidro ^[92]	2018	(2007–2015)	France	Marseille	SC	29	26	2	NR	NR	NR	NR
Ding ^[93]	2018	(2011–2016)	China	Guangzhou	SC	30.8	16	1	51.3	64	12	1
Nozdrzykowska ^[94]	2015	(2002–2013)	Germany	Leipzig	SC	NR	129	1	NR	NR	NR	NR
Lei Liu ^[95]	2016	(2013–2014)	China	Shanghai	SC	15.4	203	11	55	52.4	167	7
Hu ^[10]	2019	(2013–2017)	China	Zhejiang	SC	25.8	571	12	NR	NR	NR	NR
Gao ^[9]	2019	(2001–2013)	China	Beijing	SC	77.7	751	4	52.8	NR	619	NR

IMC = international multicenter, NMC = national multicenter, NR = not reported, RTAD = retrograde type A dissection, SC = single center, TEVAR = thoracic endovascular aortic repair.

CI: 1.9–2.8), with an I^2 heterogeneity of 44.09% ($P < .001$) (Fig. 2). The incidence of RTAD in the studies conducted in Europe (64/1718 cases; 3.7%), Asia (94/5280 cases; 1.7%), North America (81/2294 cases; 3.5%) as well as multi-continental studies (42/1266 cases; 3.3%) were similar; one smaller study in South America had a higher incidence (4/42 cases; 9.5%). With the exception of one study in South America, no significant difference was found in RTAD incidence among the continents using the Kruskal–Wallis test ($P = .08$).

Of the overall 285 cases with RTAD, time to occurrence after TEVAR was reported in 147: 89 (60.6%) occurred within 30 days; 43 (29.2%) between 1 and 12 months; 15 (10.2%) later than 1 year. Of the 89 early RTADs (within 30 days), 50 (34.0%) were intraoperative or perioperative (within 15 days of TEVAR) (see Table S4, Supplemental Digital Content, <http://links.lww.com/MD/I481>, which illustrates time to occurrence of RTAD). The Friedman test showed that the incidence of RTAD was significantly different in these time periods ($P = .005$). From the enrolled trials, 51 studies with 5058 total cases and 143 RTAD patients reported early RTAD in 94 cases (65.7%). However, 27 studies (5542 total cases and 142 RTAD patients) did not mention any information about the early occurrence of RTAD (Table 4). Forty-seven studies comprising 4592 cases and 128 RTAD patients showed late RTAD in 46 cases (35.9%). However, 31 studies (6008 total and 157 RTAD cases) did not report any information about late RTAD (Table 4). Using Wilcoxon signed-rank test, a significant difference was found in the incidence of RTAD between early and late occurrence ($P < .001$), i.e., the incidence of early RTAD was 1.8 times higher than that of late RTAD.

RTAD occurred in 2.2% (114/5230), and 0.9% (45/5169) of the cases in the acute TBAD and chronic TBAD groups, respectively (see Table S4, Supplemental Digital Content, <http://links.lww.com/MD/I481>, which illustrates time to occurrence of RTAD). Using Wilcoxon signed-rank test revealed that the proportion of RTAD patients with acute TBAD was significantly higher than those with chronic TBAD among all reported RTAD cases ($P = .008$). Twenty-four studies with 3521 patients provided comparative information on two arms of both acute

and chronic TBAD for meta-analysis. The incidence of RTAD in patients with acute TBAD was higher but not statistically significant as compared to the patients with chronic TBAD with a RR of 1.42 (95% CI: 0.95–2.12; $P = .08$; Fig. 3) using a random model. There was no heterogeneity among the studies ($P = .8$, $Chi^2 = 16.9$, and $I^2 = 0\%$).

Although 44 studies described oversizing in TEVAR, most of them provided interval ranges without a detailed numerical description. Of them, stent-graft oversizing was $\leq 10\%$ in 22 studies with 3013 TBAD cases, while it was between 10% to 20% in 20 studies with 2867 TBAD patients, and $\geq 20\%$ in only 2 studies with 350 cases (see Table S5, Supplemental Digital Content, <http://links.lww.com/MD/I482>, which illustrates stent-graft oversizing). The incidence of RTAD was 3.6% (110/3013), 2.3% (65/2867), and 3.4% (12/350) in the stent-graft oversizing categories of $\leq 10\%$, 10% to 20%, and $\geq 20\%$.

The incidence of RTAD was 2.1% (112/5328) and 0.9% (39/4381) in the proximal bare stents and non-bare stents groups, respectively (Table 2). According to Wilcoxon signed-rank test, among all reported RTAD cases, the proportion of RTAD patients in proximal bare stents group (112/153 cases; 73.2%) was not significantly different from non-bare stents group (39/129 cases; 30.2%) ($P = .11$). Fourteen studies with 2347 patients provided comparative information on two arms of both proximal bare and non-bare stents for meta-analysis. Pooled meta-analysis showed that the incidence of RTAD in proximal bare stents group was 2.1-fold higher than non-bare stents group with a RR of 1.55 (95% CI: 0.87–2.75; $P = .13$; Fig. 4) using a random model. There was no heterogeneity among the studies in this meta-analysis ($P = .71$, $Chi^2 = 9.80$, and $I^2 = 0\%$; Fig. 4).

Of 78 selected studies, 14 reported clinical manifestations of RTAD. Chest pain and sudden fluctuations in blood pressure were the main symptoms of RTAD. Four studies described RTAD as asymptomatic after TEVAR. Detailed description of the clinical presentation of RTAD is provided in Table S6, Supplemental Digital Content, <http://links.lww.com/MD/I483>, which illustrates clinical manifestation of RTAD. Of 285 cases with RTAD, 160 (56.1%) and 29 (10.2%) were

Table 2**Risk factors in type B aortic dissection patients who underwent thoracic endovascular aortic repair and those who experienced retrograde type A dissection.**

Risk factors	Studies (n)	Total TBAD patients (N)	Patients with risk factor (n)	Patients with risk factor (%)	
All patients who underwent thoracic endovascular aortic repair					
Male gender	52	7110	5534	77.8	
Hypertension	47	6134	5118	83.4	
Diabetes mellitus	37	4779	474	9.9	
Coronary artery disease	38	4477	668	14.9	
Renal impairment	38	3581	446	12.4	
Pulmonary disease	35	3369	446	13.2	
Marfan syndrome	20	2925	44	1.5	
ASA I	3	958	42	4.3	
ASA II	11	1482	279	18.8	
ASA III	13	1558	565	36.2	
ASA IV	13	1558	598	38.3	
ASA V	8	1126	52	4.6	
Smoking	34	5283	2521	47.7	
Age (yr)	57.4			NR	
Risk factors	Studies (n)	Patients with risk factor (n)	Total RTAD (n)	Total TBAD patients (N)	Total TBAD patients (%)
Patients with retrograde type A dissection					
Male gender	16	66	77	2747	85.7
Hypertension	7	44	51	2099	86.2
Diabetes mellitus	4	5	35	1634	14.2
Coronary artery disease	4	4	32	1306	12.5
Renal impairment	5	6	41	1725	14.6
Pulmonary disease	5	7	41	1725	17.0
Marfan syndrome	11	9	58	2444	15.5
ASA	NR	NR	NR	NR	NR
Smoking	4	21	32	1306	65.6
Age (yr)	56.6				NR

ASA = American Society of Anesthesiology physical status classification, NR = not reported, RTAD = retrograde type A dissection, TBAD = type B aortic dissection.

diagnosed by CT at regular and irregular imaging follow-up period, respectively. According to Kruskal Wallis Test, cumulative incidence of RTAD did not statistically differ among the studies with regular or irregular imaging follow-up period ($P = .63$). Twenty-three studies with 4412 TBAD cases and 96 RTAD patients (33.7%) did not share detailed information on imaging follow-up time (see Table S7, Supplemental Digital Content, <http://links.lww.com/MD/I484>, which illustrates imaging follow-up).

Table 5 shows the surgical and non-surgical treatment of RTAD; 52 studies comprising 7546 TBAD and 214 RTAD cases reported that 156 (72.9%) were treated surgically (Fig. 5). Eight cases (5.1%) were re-operated using the frozen elephant trunk technique. Other total arch repair (including ascending aorta repair and aortic arch repair) was performed in 16.7% (26/156); hemiarch repair or ascending aorta repair alone or Bentall procedure was performed in 19.9% (31/156); and repeated endovascular treatment was performed in 3.8% (6/156). The details of surgical approaches in 61 cases (39.1%) were not reported (see Table S8, Supplemental Digital Content, <http://links.lww.com/MD/I485>, which illustrates reported treatments of RTAD of enrolled studies). Of 9 studies comprising 73 RTAD cases and 2123 TBAD patients, 17 RTAD patients (23.2%) received non-surgical therapy including conservative wait-and-see and medical treatment (see Table S8, Supplemental Digital Content, <http://links.lww.com/MD/I485>, which illustrates reported treatments of RTAD of enrolled studies).

Death among RTAD cases was reported in 76 out of 198 RTAD cases in 52 studies with different follow-up periods (Table 6) and (see Table S9, Supplemental Digital Content, <http://links.lww.com/MD/I486>, which illustrates time and reasons of mortality of RTAD). Single arm meta-analysis estimated a mortality rate of 42.2% (95% CI: 32.5–51.8), with an I^2 heterogeneity of 70.11% ($P < .001$) (Fig. 6).

From 79 RTAD cases who died after TEVAR, the time of death was reported in 39 cases. Of whom, 24 cases (61.5%) died

within the first 30 days, 7 (17.9%) died between 1–12 months after TEVAR, and 8 (10.2%) deaths occurred one year after TEVAR. From 24 RTAD cases who died in early first month, the time of death was intraoperatively until first two weeks after TEVAR in 19 cases (20.5%). The rate of early mortality was 25 out of total 109 RTAD cases (22.9%) in 39 studies. The rate of late mortality was 11 out of a total 104 RTAD cases (10.5%) from 36 studies. However, 39 (176 cases with RTAD) and 42 studies (181 cases with RTAD) did not report any information about the early and late mortality rate of RTAD, respectively (Table 6) and (see Table S9, Supplemental Digital Content, <http://links.lww.com/MD/I486>, which illustrates time and reasons of mortality of RTAD). The rate of early mortality of RTAD was 2.1 times higher than that of late mortality. Using Wilcoxon signed-rank Test, no significant difference was found in RTAD incidence between early and late mortality of RTAD ($P = .44$).

4. Discussion

During the past decade, TEVAR has become one of the most common surgical procedure in many thoracic aortic pathologies.^[14,96–99] This method is less invasive than open surgery but still has several complications, including some new ones that are only now being characterized and understood. Some recognized complications that can occur after TEVAR include aneurysm development, aortic rupture, stroke, bowel infarction, paraplegia, limb ischemia, endoleak, and access-related complications.^[14,98,100,101] There are also important device-related complications such as stent-graft induced aortic wall injury incurred by TBAD patients after TEVAR, which can require secondary intervention if distal but can be fatal if proximal (RTAD). Although the risk of proximal SINE is low, the fatality of this complication requires vigilance in patients who develop new onset symptoms in the early period after TEVAR treatment. Careful technique, minimal oversizing, and use of disease

Table 3
Details of published reports of thoracic endovascular aortic repair and incidence of retrograde type A dissection.

First author	Year	Stent-graft detail	Total TEVARs (N)	TEVAR device			TEVAR in patients with RTAD	
				Bare stent	Non-bare stent	RTAD (n)	Bare stent	Non-bare stent
Czermak	2002	Talent (Medtronic)	5	5	0	1	1	0
Kato	2002	Z stents covered with expanded polytetrafluoroethylene (Impra); Z stents covered with woven polyester	28	0	28	1	0	1
Palmer	2002	Thoracic Excluder (Gore); Talent (Medtronic)	14	3	11	2	1	NR
Fattori	2003	Talent (Medtronic); Thoracic Excluder (Gore)	22	NR	NR	2	NR	NR
Grabenwoger	2004	Talent (Medtronic)	20	20	0	1	1	0
Hansen	2004	AneuRx (Medtronic); Talent (Medtronic); and Excluder (Gore)	24	NR	NR	1	NR	NR
Lee	2004	Custom-designed stent-grafts (Impra); 2-component system consisted of a 3-part unsupported nitinol wire stents covered with a graft of synthetic polyester fabric (Dacron; Ube)	37	NR	NR	1	1	0
Dong Xu	2005	TALENT (Medtronic); ENDOFIT (Endomed); VASOFLOW (Vascor); AEGIS (Microport); KINPRIDE (Grikin)	24	NR	NR	3	NR	NR
Fattori	2006	Talent (Medtronic)	180	180	0	2	2	0
Duebener	2007	Talent and Valiant (Medtronic)	13	13	0	1	1	0
Zipfel	2007	Talent (Medtronic), E-vita (Jotec), Zenith TX1 (Cook), Relay (Bolton Medical), Endofit (Endomed), Valiant (Medtronic), and TAG (Gore).	57	NR	NR	1	0	1
Kpodonu	2008	TAG (Gore)	91	0	91	6	0	6
Neuhauser	2008	Thoracic Excluder (Gore); Talent (Medtronic)	28	NR	NR	5	4	1
Dong	2009	Talent (Medtronic)	443	401	42	11	11	0
Chiesa	2011	Not reported	188	NR	NR	3	NR	NR
Kim	2011	Talent or Valiant (Medtronic)	41	41	0	3	3	0
Oberhuber	2011	TAG/cTAG (Gore); Captivia and Valiant (Medtronic); Zenith (Cook)	19	10	9	1	0	1
Parsa	2011	TAG (Gore), Zenith TX2 (Cook), Talent (Medtronic)	51	1	50	2	0	2
Wiedemann	2013	Talent (Medtronic); Thoracic Excluder (Gore); Relay (Bolton Medical); Endomed (LeMaitre Vascular), Cook	80	52	28	3	3	0
Lotfi	2013	TAG; 8 C-TAG (Gore); TX2; 4 TX1 (Cook); Talent; Valiant (Medtronic); Relay (Bolton Medical); Endofit (LeMaitre)	11	NR	NR	3	NR	NR
Wiedemann	2014	Talent; Thoracic Excluder; Relay; Zenith; Hemashield; Valiant	110	53	57	6	3	3
Faure	2014	Thoracic Excluder and C-TAG (Gore); Talent and Valiant (Medtronic); Zenith TX2 (Cook)	41	9	32	1	NR	NR
Idrees	2014	Gore, Cook, Medtronic	766	NR	NR	15	NR	NR
Zhang	2014	Hercules (Microport); Talent and Valiant (Medtronic); Zenith (Cook); Relay (Bolton Medical)	252	NR	NR	2	NR	NR
Gorlitzer	2012	Valiant (Medtronic); Thoracic Excluder (Gore)	29	24	5	4	4	0
Huang	2013	Talent (Medtronic); Hercules (Microport); Zenith TX2 (Cook)	563	420	143	4	4	0
Cochernec	2013	Cook; Medtronic; Gore; Relay	17	7	10	4	2	2
Shuyang Lu	2012	Talent; Valiant; Hercules; Zenith TX2	419	NR	NR	9	6	3
Yang	2012	Zenith TX2 (Cook)	61	0	61	1	0	1
Bunger	2013	Valiant (Medtronic); Zenith TX2 (Cook); Relay (Bolton Medical)	45	NR	NR	1	NR	NR
Canaud	2014	Talent, Valiant, AneuRx (Medtronic); Vasoflow (Weike Medical); Relay (Bolton Medical); Grikin (Grikin); Ankura (Lifetech); E-vita (Jotec); TAG (Gore)	309	NR	NR	11	11	NR
Lombardi	2012	Zenith TX2 (Cook)	40	0	40	3	0	3
Jia	2013	Valiant (Medtronic); Zenith TX2 (Cook); Hercules (Microport)	208	NR	NR	3	NR	NR
Li	2014	Talent (Medtronic), Relay (Bolton), Zenith TX2 (Cook), Hercules (Microport), TAG (Gore), Valiant (Medtronic)	669	168	501	6	5	1
Hanna	2014	TAG/cTAG (Gore); Zenith TX2 (Cook); Talent and Valiant with Captivia (Medtronic)	50	17	33	1	NR	NR
De Rango	2014	Zenith (Cook); TAG/cTAG (Gore); Relay (Bolton Medical); Talent and Valiant (Medtronic)	104	NR	NR	4	NR	NR
Appoo	2015	TAG and cTAG (Gore); Zenith TX2 (Cook)	16	2	14	0	0	0
Desai	2015	Valiant Captivia (Medtronic) (2 of 50; 5% at 1 year) and cTAG (Gore) (5 of 50; 10% at 1 year)	132	NR	NR	9	NR	NR
Kische	2015	Zenith; Valiant; Talent	35	NR	NR	1	NR	NR
Bockler	2016	cTAG (Gore)	24	24	0	1	1	0
Faure	2016	Excluder (Gore); TAG (Gore); Talent (Medtronic); Valiant (Medtronic), and Zenith TX2 (Cook)	33	NR	NR	1	NR	NR
Wang	2016	Talent (Medtronic); Captivia (Medtronic); Zenith TX2 (Cook); TAG (Gore); (Microport)	360	NR	NR	5	4	NR
Asaloumidis	2017	Talent (14); TAG (13); Excluder (2); Valiant (2); Captivia (6); Relay (2); AneuRx (1)	40	24	16	2	2	0

(Continued)

Table 3
(Continued)

First author	Year	Stent-graft detail	Total TEVARs (N)	TEVAR device			TEVAR in patients with RTAD	
				Bare stent	Non-bare stent	RTAD (n)	Bare stent	Non-bare stent
Zhao Liu	2017	Talent and Captivia (Medtronic); TX- 1/TX-2 (Cook); Hercules (Microport); Sinus (OptiMed)	58	NR	NR	6	NR	NR
Min-Hong Zhang	2017	Not reported	85	NR	NR	3	NR	NR
Tjaden	2018	CTAG or TAG (Gore)	264	264	0	6	6	NR
Tao Ma	2018	Talent and Valiant (Medtronic), Zenith TX2 (Cook), Hercules and Castor (Microport), Ankura (Lifotech), Relay (Bolton Medical), EndoFit (LeMaitre), E-vita (Jotec), and TAG (Gore).	852	NR	NR	27	NR	NR
Laquian	2018	Not reported	27	NR	NR	1	NR	NR
Chen	2018	Not reported	167	NR	NR	1	NR	NR
Piotr Buczkowski	2019	Zenith (Cook), JOTEC and Gore	68	0	68	2	0	2
Eleshra	2020	Not reported	64	NR	NR	1	NR	NR
Fukushima	2019	Zenith TX2 Pro-Form (Cook Medical), cTAG (Gore), Relay (Terumo Aortic), Najuta (Kawasumi)	24	NR	NR	0	NR	NR
Wang	2019	Valiant (Medtronic), CTAG (Gore), and TX2/Alpha (Cook Medical)	397	NR	NR	6	NR	NR
Yammine	2019	Valiant (Medtronic)	186	172	0	15	15	0
Miura	2019	Relay (Terumo Aortic)	22	22	0	0	0	0
Chassin-Trubert	2020	Valiant Captivia (Medtronic)	17	17	0	0	0	0
Pellenc	2019	TX2/TX2 alpha (Cook); cTAG (Gore); Relay (Terumo Aortic); Valiant (Medtronic)	20	NR	NR	0	0	0
Jiechang Zhu	2018	Valiant (Medtronic), Relay (Terumo Aortic) and Ankura (Lifotech)	20	20	0	0	0	0
Riesterer	2018	Relay NBS (non-bare stent) (Bolton Medical/Terumo Aortic)	34	0	34	1	0	1
Giles	2019	Not reported	258	NR	NR	12	NR	NR
Kuo	2019	TAG/cTAG (Gore), TX2/Alpha (Cook), Valiant (Medtronic)	71	40	31	2	NR	NR
Joo	2019	Valiant (using the Captivia delivery system; Medtronic), Seal (S&G Biotech), TX2 (Cook), TAG (Gore), and unidentified	17	13	2	2	2	0
Cao	2020	Zenith TX2 (Cook), Valiant (Medtronic), CTAG (Gore), Hercules (MicroPort) and Ankura (Lifotech)	76	65	11	4	3	1
El-Beyrouti	2020	RelayPro NBS (Terumo Aortic)	5	NR	NR	0	NR	NR
Charltonouw	2018	Not reported	43	NR	NR	3	NR	NR
Lou	2020	Valiant with Captivia (Medtronic), Zenith TX 2 (Cook), and CTAG (Gore)	91	80	11	3	NR	NR
Lee	2020	Seal (S&G Biotech); Valiant (Medtronic); Zenith TX2 (Cook)	87	51	36	2	NR	NR
Oshi	2020	TAG or cTAG (Gore), Valiant (Medtronic), Zenith TX2 (Cook), and Relay Plus (Terumo Aortic)	40	NR	NR	1	NR	NR
Puech-Leao	2020	Not reported	42	NR	NR	4	NR	NR
Sobocinski	2020	Not reported	41	NR	NR	2	NR	NR
Shuo Zhao	2020	Not reported	79	NR	NR	1	NR	NR
Bavaria	2015	Valiant Captivia (Medtronic)	50	50	0	2	2	0
Pedro	2018	TAG/cTAG (Gore); Valiant/Talent (Medtronic); Zenith/Pro-Form (Cook)	26	NR	NR	2	NR	NR
Ding	2018	Valiant (Medtronic); Ankura (Lifotech); ZTEG-2PT (Cook)	16	15	1	1	1	0
Nozdrzykowska	2015	TAG/cTAG (Gore); Talent/Valiant/Captivia (Medtronic); Zenith (Cook); and Endofit (LeMaitre Vascular)	129	NR	NR	1	NR	NR
Lei Liu	2016	Zenith TX2 (Cook); CTAG (Gore); Talent (Medtronic); and Hercules, Aegis, and Ankura (Microport)	203	85	118	11	5	6
Hu	2019	Valiant (Medtronic), TAG (Gore), Zenith TX2 (Cook), and Ankura (Lifotech).	571	NR	NR	12	8	4
Gao	2019	GRIMED (GRIMED) in 234 patients, Talent (Medtronic) in 20, Valiant (Medtronic) in 173, Hercules (MicroPort) in 125, Zenith TX2 (Cook) in 86, Relay (Bolton Medical) in 76 and E-vita (Jotec) in 37.	751	665	86	4	NR	NR

NR = not reported, RTAD = retrograde type A dissection, TEVAR = thoracic endovascular aortic repair.

specific stent-grafts may reduce the risk for RTAD. Distally, SINE is more frequently seen during follow-up in patients treated for chronic dissection. The most important risk factor is oversizing of the stent-graft compared to the true lumen distal landing zone.^[102] Therefore, procedure and device-related factors, the natural progression of initial aortic dissection, and unfavorable aortic-dissection anatomy are among the etiological factors mentioned.^[13,103]

The RTAD rate after TEVAR might be reduced by improving stent-graft design (non-bare stents and tapering, for example), limited oversizing, and more careful manipulation during deployment.^[13] It can also be argued that most of the information

and hypotheses about this complication are not well-cited because RTAD has been reported as a rare complication with limited information in each study. To this end, we decided to thoroughly evaluate and analyze all available information about RTAD after TEVAR in TBAD patients.

Our single-arm meta-analysis estimated that the incidence of RTAD after TEVAR in patients with TBAD to be 2.3%. Therefore, it is not a very common complication. There is probably a difference in the incidence of RTAD after TEVAR on different continents. There are also several factors affecting it, such as the genetic background of connective tissue diseases, stents that have been used before, and differences in procedure-related

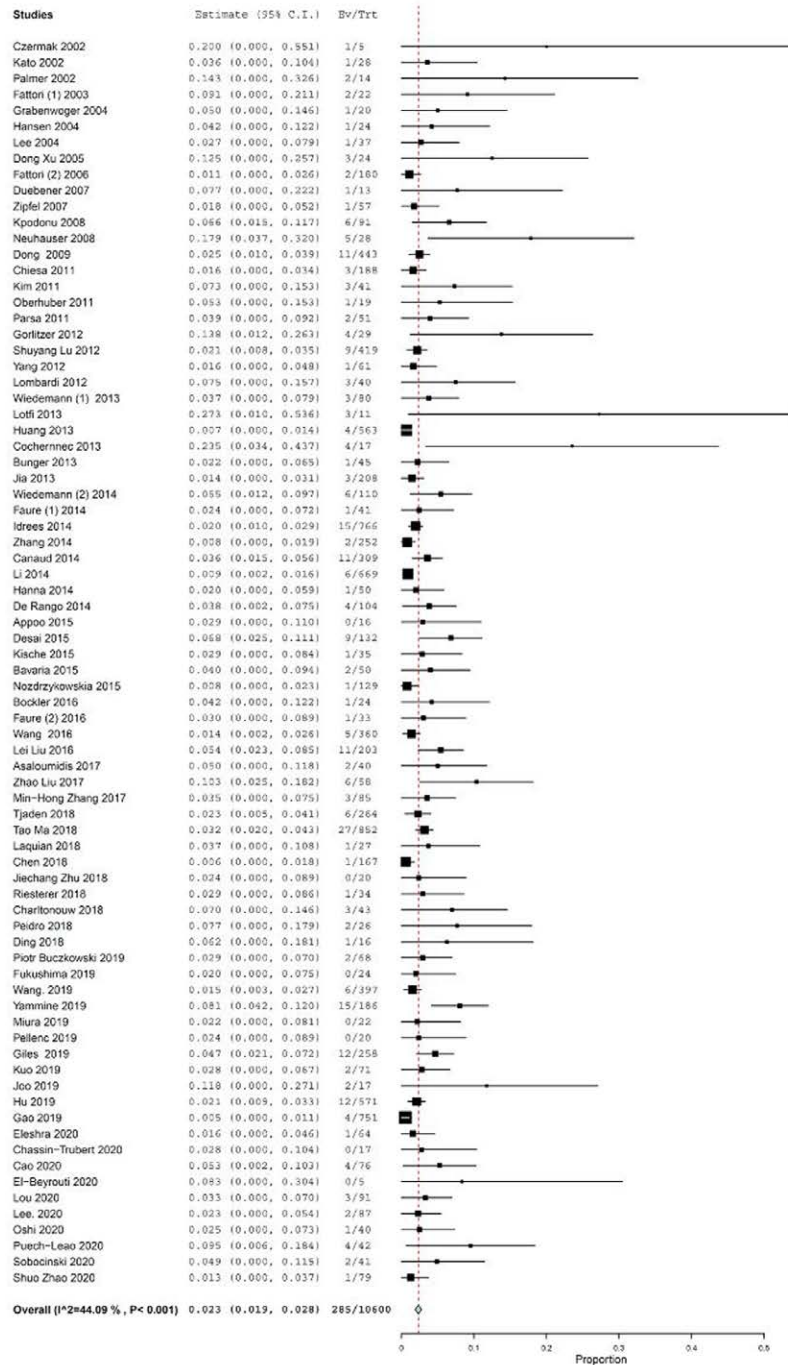


Figure 2. Forest plot of proportion single-arm meta-analysis for RTAD after TEVAR. RTAD = retrograde type A dissection, TEVAR = thoracic endovascular repair.

factors. However, it cannot be ignored that the incidence of RTAD has been less pronounced in Asian studies. On the other hand, most Asian studies have been conducted in China. Besides, the incidence of RTAD is similar on the continents of America and Europe and higher than the reported incidences in Asian studies. Consequently, although this complication is considered rare, it needs to be greater attention in European and American countries. Centers with < 50 TBAD cases undergoing TEVAR were 2.26 times more likely to incur RTAD compared to centers with > 50 TBAD cases. As a result, it can be acknowledged that highly experienced centers reported a lower incidence of RTAD, suggesting the important hypothesis that this complication was significantly related to the procedure and postoperative management, strongly dependent on the surgeon's

experience. The decline in RTAD incidence from the introduction of TEVAR to the present may support the hypothesis that the incidence of RTAD decreases with increased experience and better technique. In general, it may be concluded that in China, due to the large population and existence of certain TEVAR centers with a certain number of surgeons, the surgeons have probably more experience in performing TEVAR. European and American countries, while being less populated, have more centers performing TEVAR. For this reason, most surgeons may not yet have reached their full potential. For instance, the risk of RTAD occurrence can increase when surgeons pass a guide wire through a tortuous aortic arch. The risk is exacerbated when getting it through anatomically abnormal areas or when the aorta is distorted or very thin, meaning that any friction

Table 4
Timing of RTAD.

Time post-TEVAR	Studies (n)	Patients (n)	RTAD (n)	TEVAR (n)
0–14 d	46	50	128	3730
Early (within 30 d)	50	89	153	4834
1–12 mo	46	43	138	4368
After 1 yr	47	15	141	4556
Early RTAD				
Reported	51	94	143	5058
Not reported	27	Not reported	142	5542
Late RTAD				
Reported	47	46	128	4592
Not reported	31	Not reported	157	6008

RTAD = retrograde type A dissection, TEVAR = thoracic endovascular aortic repair.

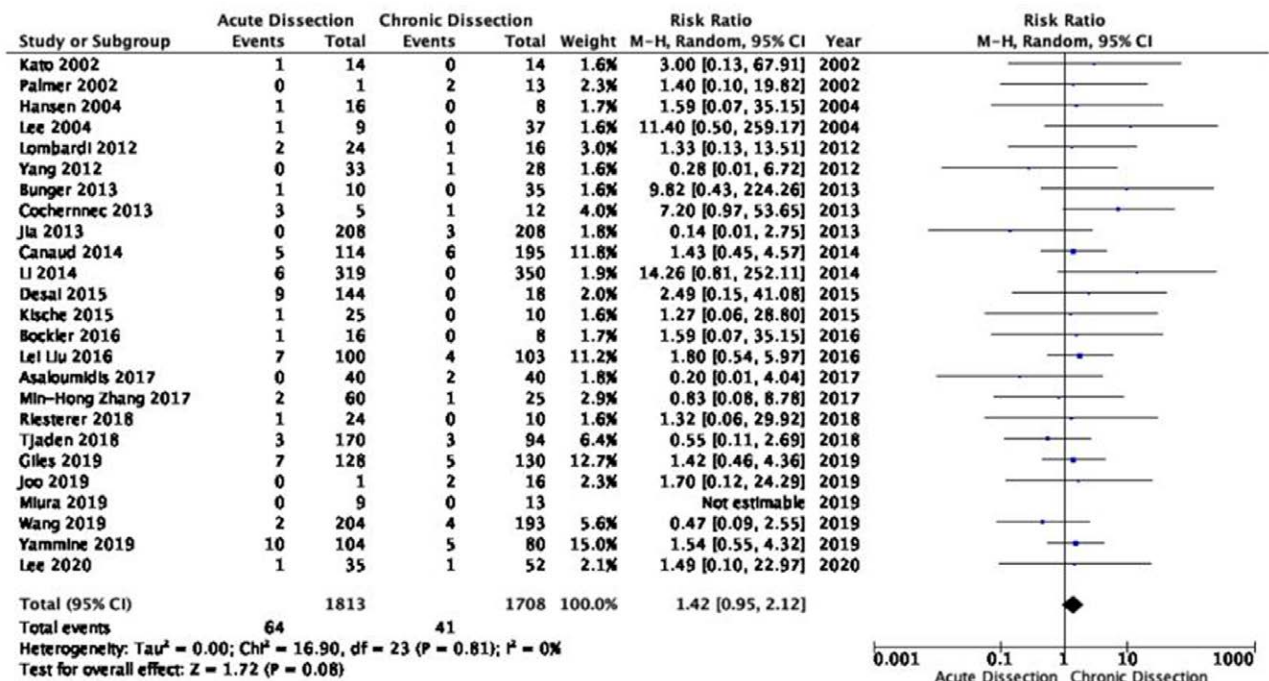


Figure 3. Forest plot for comparing of rates of RTAD post-TEVAR between acute and chronic type of TBAD. CI = confidence interval, RTAD = retrograde type A dissection, TBAD = type B aortic dissection, TEVAR = thoracic endovascular repair.

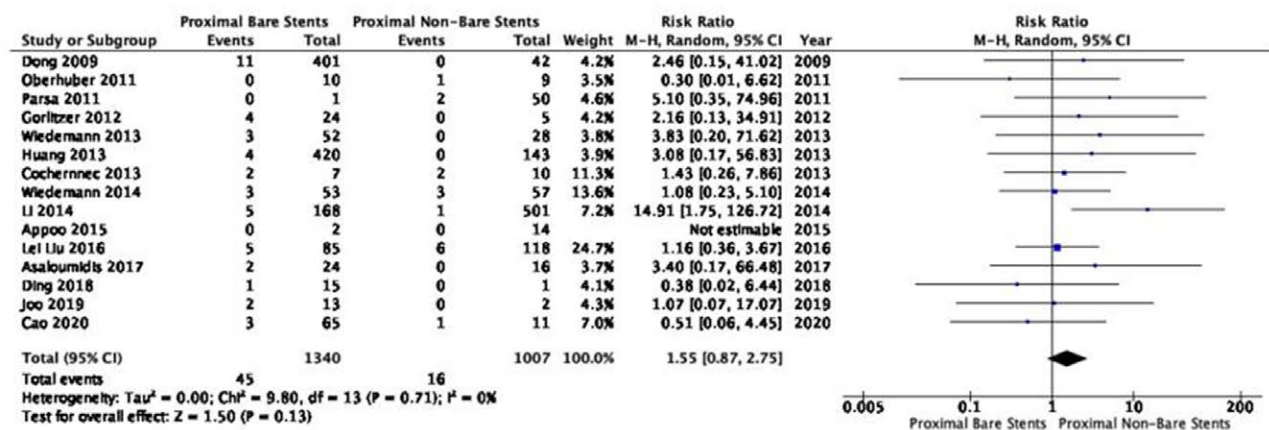


Figure 4. Forest plot for comparing of rates of RTAD post-TEVAR between implanted proximal bare and non-bare stents. CI = confidence interval, RTAD = retrograde type A dissection, TEVAR = thoracic endovascular repair.

Table 5
Therapeutic options of RTAD.

	Treatment	Nr of studies	Nr of treatment	Total RTAD	Total TEVAR
A) Therapeutic options					
Non-surgical	Reported	9	17	73	2123
	ND	69	ND	212	8477
Surgical	Reported	52	156	214	7546
	ND	26	ND	71	3054
Interventions					
B) Surgical interventions					
Surgical treatment	No exact data about open repair	22	61	39.10	
	Total arch repair (Ascending Aorta + aortic arch replacement)	12	26	16.67	
	Ascending repair or hemiarch repair or Bentall procedure or aortic root	22	31	19.87	
	Ascending TEVAR or Re-Stent or Stent-Dilatation	4	6	3.85	
	Frozen Elephant Trunk	2	8	5.13	
	* Undifferentiated	2	24	15.38	

RTAD = retrograde type A dissection, TEVAR = thoracic endovascular aortic repair.

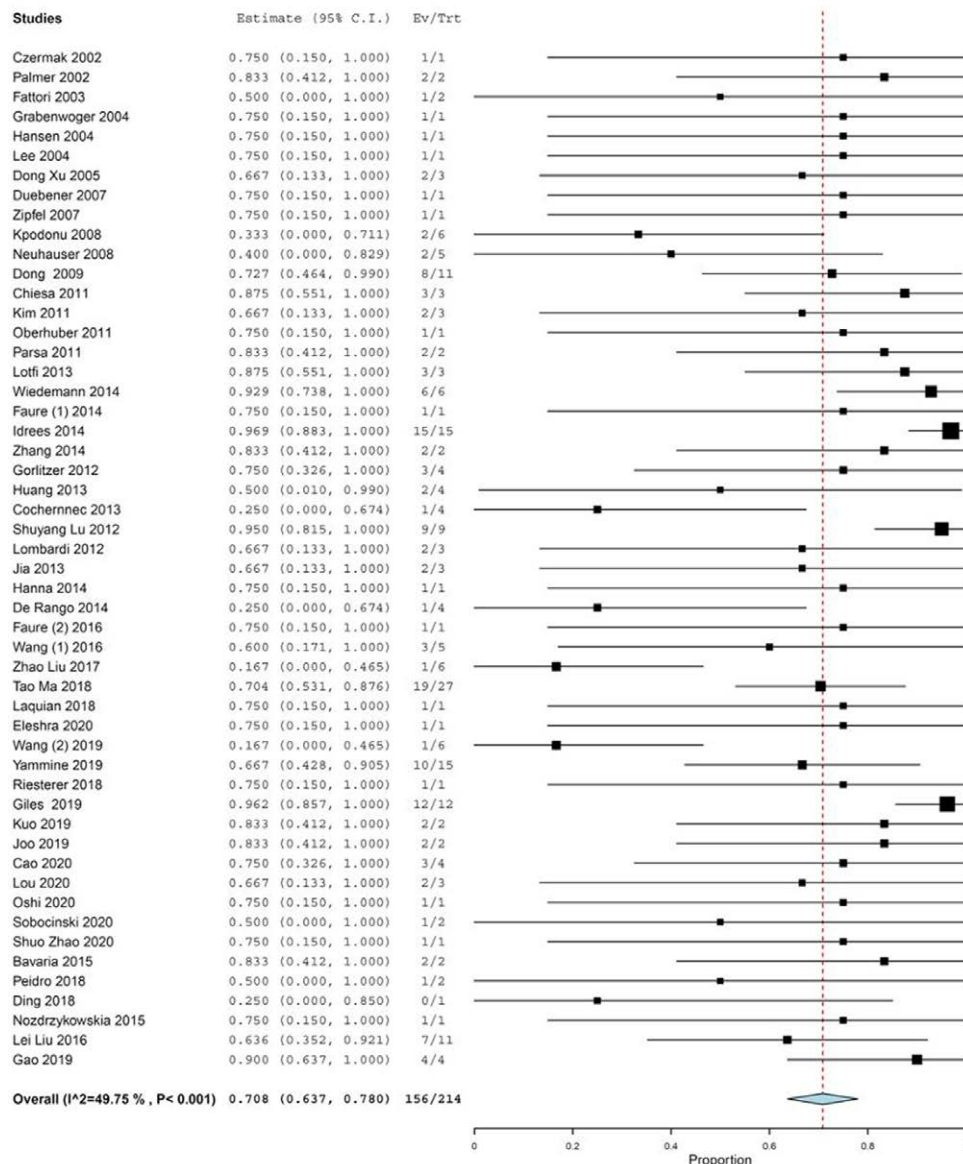


Figure 5. Forest plot of proportion single-arm meta-analysis for surgical re-intervention as therapeutic option of RTAD after TEVAR. CI = confidence interval, RTAD = retrograde type A dissection, TEVAR = thoracic endovascular repair.

from catheter or guide wire can damage the wall. Such risks can be effectively mitigated by more experienced centers and surgeons.^[13,16]

Our findings also showed that RTAD occurred primarily as a hyperacute or acute condition rather than a chronic condition. Thus, the first month after TEVAR was the maximum duration for RTAD incidence; in addition, from the moment of TEVAR operation until the first two weeks, the probability of its occurrence was the highest. Our estimates revealed that the incidence of early RTAD was approximately 1.8 times higher than that of late RTAD. One hypothesis is that patients with acute TBAD are more likely to have urgent or emergent TEVAR which may be less accurate in preoperative assessments compared with chronic TBAD patients. Moreover, acute pathological changes in the aorta may increase the probability of extension of dissection and therefore predispose to RTAD. Having said that, Tjaden et al^[67] found no significant difference between the risk of RTAD in acute compared to chronic TBAD. In this meta-analysis, the number of RTAD patients with acute TBAD was significantly higher than the number of RTAD patients with chronic TBAD. However, the corresponding risk ratio of 1.42 was not statistically significant. Although the findings were borderline, clinically, it can be accepted that RTAD could be more incurred by patients with acute TBAD. Therefore, more accurate diagnostic and therapeutic evaluation should be adopted to prevent this complication in acute TBAD cases. After evaluating the data, it was shown that there were no significant regional differences in the availability of follow-up data and imaging data.

Some other studies reported that proximal bare stent configuration was associated with an increased risk of RTAD.^[104] Chen et al claimed that with a risk ratio of 2.06, the incidence of RTAD in TEVAR was higher in the proximal bare stent than the proximal non-bare stent.^[103] This meta-analysis found that risk of RTAD in the proximal bare stents group was 2.1-fold more than in the proximal non-bare stents group. According to our comparative meta-analysis, the difference in the incidence of RTAD was not significant in the two groups of proximal bare stents and non-bare stents with a risk ratio of 1.50. This finding can be interpreted in the way that the quality of proximal bare stents design and the experience of surgeons working with these stents' models have probably increased in recent years. However, it cannot be ignored that according to previous studies, the percentage of RTAD in the proximal bare stent group was higher, even though it was not significant. Besides, it is clinically significant that if the patient is at risk of RTAD after TEVAR, such as patients with Marfan syndrome, connective tissue diseases, and acute TBAD who want

to undergo non-elective TEVAR, proximal non-bare stents might be the best choice.

Dong et al explained that using angiotensin-converting enzyme inhibitors, B-blockers, calcium antagonists, or angiotensin receptor blockers was only suggested as medical management procedures when RTAD was limited, and the patient's situation was clinically stable.^[105] In the present study, 11.5% of the studies with 73 patients reported non-surgical treatment with RTAD, implying that conservative wait-and-see treatment or re-surgical treatment was not accepted by patients, hence the use of non-surgical treatments. It is clear that surgical treatment should be applied in patients with unstable and limited progression since using drug treatment is not sufficient. Our findings suggested that some of the most common surgical reinterventions could treat RTAD, including ascending aorta repair alone, hemi-arch replacement, and Bentall procedure. Clinically, after RTAD diagnosis followed by TEVAR, it is recommended to make treatment decisions by an interdisciplinary aortic team including vascular surgeons, cardiac surgeons, radiologists, intensive-care specialists, and anesthesiologists to evaluate the re-intervention carefully and to manage clinical and radiological follow-ups and postoperative care.

The RTAD mortality rate post-TEVAR, although low, was significantly higher than spontaneous type A aortic dissection.^[106,107] This was clearly more significant during the first month post-TEVAR compared to 1 to 12 months, and one year after TEVAR. Of those who died due to RTAD during the first month after TEVAR, 79.1% died during surgery or in the first hours and days after surgery. Due to the significant and high mortality rate of this uncommon complication, RTAD should be considered as one of the differential diagnoses with high risk during ICU stay or hospital stay after surgery and even after discharge. If the patient suddenly suffers from any chest pain, back pain, chest discomfort, sudden changes in blood pressure, syncope, or any other sudden clinical signs, appropriate radiological evaluations should be performed to perform appropriate reintervention as soon as possible and to avoid sudden death. Numerous studies have also suggested that the occurrence of RTAD coincides with the onset of multi-organ failure and eventual death.^[108,109] It should be mentioned that most research done on RTAD had a small sample size, and the mortality rate varied according to various treatment strategies applied.^[108,109] Hence further well-designed, large scale clinical trials with longer-term follow-up are needed to accurately evaluate mortality rate of RTAD after TEVAR and its diagnostic workout and surgical management. We recommend that future studies investigate

Table 6
Mortality of RTAD.

Situation of report		Nr of studies	Nr of dead	Total RTAD	Total TEVAR of these Studies	Mortality rate
A)Report of mortality						
Reported		52	76	198	6915	38.30%
Not reported		26	43	87	3685	
Time of mortality		Nr of studies	Nr of dead	Total RTAD	Total TEVAR	Mortality rate
B)Time interval of mortality						
0–14 d		38	19	107	3473	17.76
Early 30 d		39	24	109	3514	22.02
1–12 mo		37	7	106	3446	6.60
After 1 yr		36	8	104	3375	7.69
C)Early or late mortality						
Early mortality	Reported	39	25	109	3514	22.94
	ND	39	ND	176	7086	
Late mortality	Reported	36	11	104	3375	10.58
	ND	42	ND	181	7225	

RTAD = retrograde type A dissection, TEVAR = thoracic endovascular aortic repair.

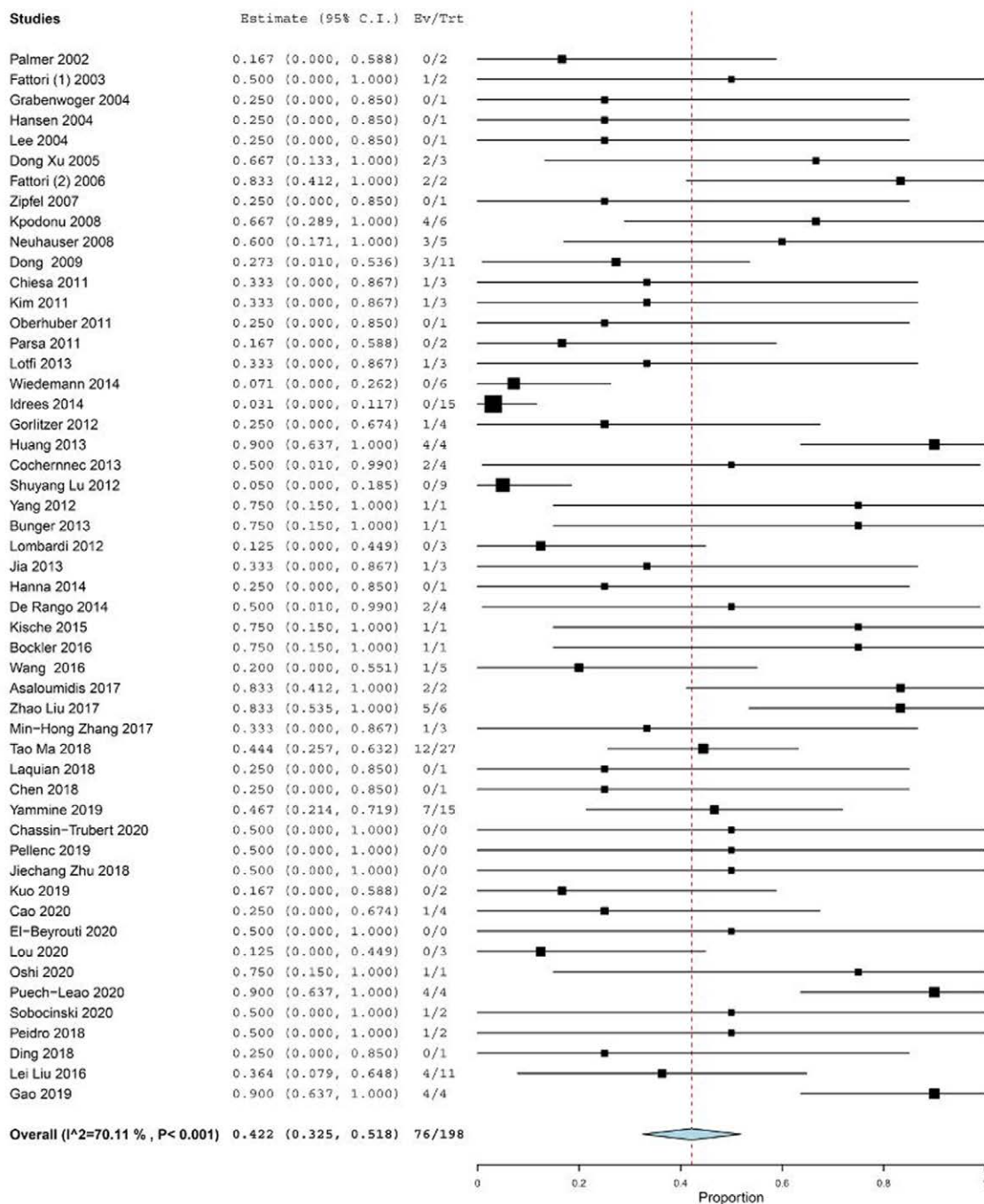


Figure 6. Forest plot of proportion single-arm meta-analysis for the mortality rate of RTAD after TEVAR. CI = confidence interval, RTAD = retrograde type A dissection, TEVAR = thoracic endovascular repair.

the correlation between genetic parameters and incidence for RTAD, as well as patients who die due to RTAD.

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