# Management of complicated and uncomplicated acute type B dissection. A systematic review and meta-analysis

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**Background:** The management of acute type B dissection represents a clinical challenge. We undertook a systematic review of the available literature regarding medical, surgical and endovascular treatments of acute type B aortic dissection and combined the eligible studies into a meta-analysis.

**Methods:** An extensive electronic health database search was performed on all articles published from January 2006 up to November 2013 describing the management of acute type B aortic dissection. Studies including less than 15 patients were excluded.

**Results:** Acute complicated type B dissection: overall, 2,531 patients were treated with endovascular repair (TEVAR) and the pooled rate for 30-day/in-hospital mortality was 7.3%. The pooled estimates for cerebrovascular events, spinal cord ischemia (SCI) and total neurologic events were 3.9%, 3.1% and 7.3%, respectively. A total of 1,276 patients underwent open surgical repair and the pooled rate for 30-day/in-hospital mortality was 19.0%. The pooled rate for cerebrovascular events was 6.8%, for SCI 3.3% and for total neurologic complications 9.8%. Acute uncomplicated type B dissection: outcome of 2,347 patients who underwent conservative medical management were analyzed. The pooled 30-day/in-hospital mortality rate was 2.4%. The pooled rate for cerebrovascular events was 1%, for SCI 0.8% and for overall neurologic complications 2%.

**Conclusions:** Endovascular repair provides a superior 30-day/in-hospital survival for acute complicated type B aortic dissection compared to surgical aortic reconstruction. However, open repair still has a significant role as endovascular repair is not applicable in all patients and there remains concerns regarding the durability of this technique. TEVAR seems to have a more favorable outcome regarding aortic remodeling and the aortic-specific survival rate when compared with medical therapy alone. Randomized controlled trials focusing on the prognostic factors of early and late complications in uncomplicated type B dissections are needed.

Keywords: Type B aortic dissection; acute; medical treatment; endovascular



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#### Introduction

Aortic dissection is a potentially life-threatening condition that occurs when a tear is formed in the wall of the aorta. Stanford type B or DeBakey III aortic dissection originates in the descending thoracic aorta without retrograde extension into the ascending aorta (1). Acute type B dissections may be classified as uncomplicated or complicated. Approximately 25% of patients presenting with acute type B aortic dissection are complicated at admission by malperfusion syndrome or hemodynamic instability, resulting in a high risk of early death if untreated (1-3). Complicated type B aortic dissection refers to malperfusion

syndrome involving visceral, renal, or extremity ischemia, rupture or impending rupture, uncontrolled hypertension, persistent abdominal or chest pain, or findings of rapid expansion on computed tomography (CT) imaging.

The management of acute complicated type B dissection represents a clinical challenge. Uncomplicated dissections have traditionally been managed non-operatively with aggressive blood pressure control. However, best medical treatment (BMT) is associated with a considerable risk of disease progression to complicated dissection or aneurysmal degeneration of the affected aortic segment, which is the most feared complication in the long term, involving about 30-40% of patients (4,5). The lower morbidity and mortality rates associated with endovascular procedures have generated interest in the treatment of uncomplicated dissections.

We undertook a systematic review of the available literature regarding medical, surgical and endovascular treatments of acute type B aortic dissection. We combined the eligible studies into a meta-analysis with the intention of studying the efficacy of these treatment strategies.

#### Methods

Definitions

#### Acute type B aortic dissection

Aortic dissection involving the descending thoracic aorta and distal sites only and the diagnosis has to be within 14 days of onset of symptoms.

#### Complicated type B aortic dissection

Complicated dissections refer to aortic rupture, visceral and renal ischemia, lower extremities ischemia, or spinal cord ischemia (SCI). Expansion to the aortic arch or proximal descending aorta with a total diameter of 4.5 cm or greater is also considered a complicated dissection. However, refractory hypertension, hypertension persisting despite three different classes of antihypertensive therapy at maximal recommended or maximal tolerated doses, if not present in the clinical history before the onset of dissection, is considered a sign of instability or of renal malperfusion.

Malperfusion syndrome is the most frequent complication of type B dissection. The clinical presentation includes paraparesis or paraplegia, lower limb ischemia, abdominal pain, nausea, or diarrhea. Visceral artery malperfusion may be associated with an increase in laboratory markers (bilirubin, amylases, hepatic and intestinal enzymes). The CT angiography or magnetic resonance angiography findings such as true lumen compression, or an intimal flap inside the renal, celiac, or mesenteric arteries, carry a high suspicion of visceral malperfusion. Delay or absence of nephrographic effect during the late phase of contrast-enhanced CT scan, often accompanied by an increase in serum creatinine and/or refractory hypertension, indicates renal malperfusion.

#### Search strategy

The present meta-analysis was conducted in accordance with the recommendations of the Meta-Analysis of Observational Studies in Epidemiology (MOOSE) group (6). An extensive electronic health database search was performed on all articles published from January 2006 up to November 2013 describing the management of acute type B aortic dissection. The search was performed using exploded medical subject heading (MeSH) terms: "acute type B aortic dissection", "complicated", "uncomplicated", "medical treatment", "surgical treatment", "open repair" and "endovascular treatment". Publications were retrieved through electronic search engines (Medline, Embase, Scopus, Google Scholar, Ovid, and the Cochrane Library). All studies were independently assessed, and full texts of potentially eligible studies were retrieved. In addition, the reference lists of all retrieved articles were examined for further relevant series.

#### Study eligibility, and exclusion criteria

Studies were included in the present review if (I) the index aortic pathology was acute type B aortic dissection; (II) BMT, open surgical repair or TEVAR were the applied treatment options; (III) stated the incidence of at least one of the basic outcome criteria; (IV) included  $\geq 15$  patients. Articles in languages other than English were eliminated from further analysis. Case reports and case series with less than 15 patients were excluded. Studies referring to chronic aortic dissection were excluded. Studies referring to type A aortic dissection or to combined hybrid endovascular and open thoracic aorta repairs were excluded as well. When multiple publications on the same patient population were identified or study populations overlapped, only the latest report was included, unless the reported outcomes were mutually exclusive. Furthermore, several studies included patients with acute type B dissection as a subset of the entire study cohort. These were included in the present review if separate data for this patient subgroup was provided.

The available data were independently extracted and

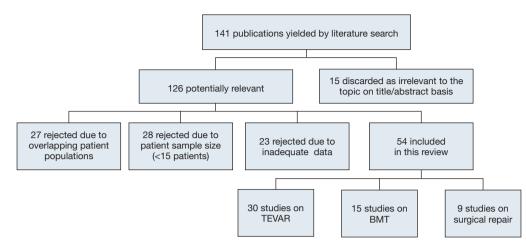


Figure 1 Study flow chart. TEVAR, treated with thoracic endovascular aortic repair; BMT, best medical treatment.

analyzed by two reviewers (S.M. and K.M.), and a consensus was reached if discrepancies were observed. Primary endpoints included peri-procedural (30-day/in-hospital) mortality, stroke, SCI and total neurologic complications rates. Data regarding long-term survival and aortic event freedom were also analyzed.

#### Statistical analysis

Separate meta-analyses was carried out on all eligible studies for peri-procedural (30-day/in-hospital) mortality, stroke, SCI and total neurologic complications. The pooled proportion was calculated as the back-transformation of the weighted mean of the transformed proportions. Statistical heterogeneity was measured using the Cochran Q statistic score (P<0.10 was considered indicative of statistically significant heterogeneity) and the  $I^2$  test. A fixed-effects model was used when no heterogeneity existed among studies. Otherwise, the random effects model was used. The meta-analyses were conducted using StatsDirect statistical software (StatsDirect Ltd, UK). Frequency study-specific estimates were pooled and are reported as event rates with corresponding 95% confidence intervals (95% CI). Longterm data were reported with Kaplan-Meier rates by the study investigators. No pooled analyses of long-term results were performed as there were variable event definitions among studies.

#### **Results**

The literature search identified 141 potentially relevant citations. Fifteen were excluded at the title or abstract level

as being irrelevant. Of the remaining 126, 27 included overlapping patient populations and were excluded, 28 reported on series of <15 patients and were also excluded, and a further 23 publications reported insufficient data on study outcomes. Eventually, 54 studies were considered eligible and included in the present meta-analysis. This included 30 studies with a total of 2,531 patients with acute type B aortic dissection treated with TEVAR, 15 studies (5,20,30,32,37-47) with a total of 2,347 patients treated with best medical therapy and 9 studies (22,24,35,38,48-52) with a total of 1,276 patients treated with open surgical repair (*Figure 1*).

#### Acute complicated type B dissection

### Endovascular treatment for acute complicated type B dissection

Overall, 2,531 patients with acute type B aortic dissection were treated with TEVAR (*Table 1*). The pooled rate for 30-day/ in-hospital mortality was 7.3% (95% CI, 5.3% to 9.6%). The pooled estimates for cerebrovascular events, SCI and total neurologic events were 3.9% (95% CI, 3.2% to 4.8%), 3.1% (95% CI, 2.0% to 4.4%) and 7.3% (95% CI, 5.2% to 9.7%), respectively (*Figure 2*). Survival rates ranged from 62% to 100% at 1-year and from 61% to 87% at 5-years, whereas freedom from aortic events ranged from 45% to 77%.

#### **Open surgical repair**

A total of 1,276 patients from nine studies who underwent open surgical repair for acute complicated type B aortic dissection were analyzed (*Table 2*). The pooled rate for

Author	Period of study	N (pts)	Indication	30-da	y/in-hos	spital (n)	FU (Months)	Survival rate (%)			Aortic event freedom rate (%)		
Author				CVE	SCI	Mortality		1 year	3 years	5 years	1 year	3 years	5 years
Di Tommaso 2006 (7)	2001- 2005	26	Acute complicated	-	0	0	29	ND	ND	ND	ND	ND	ND
Chen 2006 (8)	2001- 2005	23	Acute complicated	1	0	1	27.5	ND	ND	ND	ND	ND	ND
Yang 2006 (9)	2001- 2005	36	Acute complicated	0	0	1	15	ND	ND	ND	ND	ND	ND
Jing 2008 (10)	2002- 2007	32	Acute complicated	0	0	1	18	ND	ND	86.4	ND	ND	73.9
Sayer 2008 (11)	2000- 2007	38	Acute complicated	2	0	1	30	ND	93.0	ND	ND	ND	ND
Rodriguez 2008 (12)	2000- 2006	59	Acute complicated	3	3	1	15.6	ND	ND	ND	ND	ND	ND
Böckler 2009 (13)	1997- 2008	23	Acute complicated	0	0	6	24	62	62	62.0	64	ND	45
Alves 2009 (14)	1997- 2004	45	Acute complicated	-	-	3	35.9	ND	ND	ND	ND	ND	ND
Conrad 2009 (15)	2005- 2007	33	Acute complicated	4	2	4	12	ND	ND	ND	ND	ND	ND
Guangqi 2009 (16)	2001- 2006	72	Acute complicated	3	0	1	14.4	98.6	75.0	ND	51.4	ND	ND
Feezor 2009 (17)	2005- 2007	33	Acute complicated	4	5	7	5	ND	ND	ND	ND	ND	ND
Manning 2009 (18)	2001- 2008	45	Acute complicated	2	4	5	30	ND	ND	ND	ND	ND	ND
Sze 2009 (19)	2000- 2007	23	Acute complicated	2	1	4	22.3	ND	ND	ND	ND	ND	ND
Chemelli- Steingruber 2010 (20)	1996- 2008	38	Acute complicated	1	0	5	33	81.5	ND	69.0	ND	4 retrograde type A- open repair	ND r
Botsios 2010 (21)	2001- 2006	32	Acute complicated	ND	1	3	32.1	ND	ND	ND	ND	ND	ND
Zeeshan 2010 (22)	2002- 2010	45	Acute complicated	3	6	2	37	82.0	79.0	79.0	ND	ND	ND
Torsello 2010 (23)	2005- 2008	32	Acute complicated	0	0	0	23.1	ND	ND	ND	ND	ND	ND
Brunt 2011 (24)	2005- 2008	991	Acute complicated	37	32	107	ND	ND	ND	ND	ND	ND	ND
Tang 2011 (25)	2007- 2008	30	Acute complicated	ND	0	1	12	ND	ND	ND	ND	ND	ND
O'Donnell 2011 (26)	2005- 2008	28	27 acute complicated	3	1	2	21	ND	ND	ND	ND	ND	ND

A settle set	Period	N	Indication	30-day	//in-hos	pital (n)	FU (Months)	Survival rate (%)			Aortic event freedom rate (%)		
Author	of study	(pts)		CVE	SCI	Mortality		1 year	3 years	5 years	1 year	3 years	5 years
Steuer 2011 (27)	1999- 2009	50	50 acute complicated, 10 subacute complicated	3	1	2	44.4	ND	90.0	87.0	ND	68. 0	65.0
Thomson 2011 (28)	2006- 2009	50	Acute complicated	4	1	4	23.1	ND	ND	ND	ND	ND	ND
Zipfel 2011 (29)	2006- 2008	25	Acute complicated	ND	ND	5	ND	ND	ND	ND	ND	ND	ND
Fattori 2013 (30)	1995- 2012	276	Acute (163 rupture/ malperfusion)	6	3	30	60	91.9	76.2	75.0	79.6	77.1	69.4
Ehrlich 2013 (31)	1998- 2004	29	Acute complicated	2	0	5	53	79.0	ND	61.0	82.0	ND	77.0
Qin 2013 (32)	2004- 2008	152	Acute (137 complicated 15 uncomplicated)	2	2	3	45.3	100	93.0	66.0	97.0	89.0	67.0
Shu 2013 (33)	2000- 2009	127	Acute complicated	ND	ND	2	19.1	ND	ND	ND	ND	ND	ND
Hanna 2013 (34)	2005- 2012	50	Acute complicated	1	1	0	33.8	ND	ND	84.0	76.0	ND	ND
Wilkinson 2013 (35)	1995- 2012	36	Acute complicated	3	2	4	27.7	ND	ND	ND	ND	ND	ND
Sobocinski 2013 (36)	2004- 2011	52	Acute complicated	1	5	5	25	90.4	ND	ND	ND	ND	ND

CVE, cerebrovascular event; SCI, spinal cord ischemia; ND, not determined; FU, follow-up.

30-day/in-hospital mortality was 19.0% (95% CI, 16.8% to 21.1%, *Figure 3A*). The pooled rate for cerebrovascular events was 6.8% (95% CI, 5.4% to 8.2%), for SCI 3.3% (95% CI, 2.4% to 4.4%) and for total neurologic complications 9.8% (95% CI, 8.2% to 11.5%) (*Figure 3B-D*). Survival rates ranged from 74.1% to 86.0% at 1-year and from 44.0% to 82.6% at 5-years, whereas freedom from aortic events could not be estimated as there were no available data.

#### Acute uncomplicated type B dissection

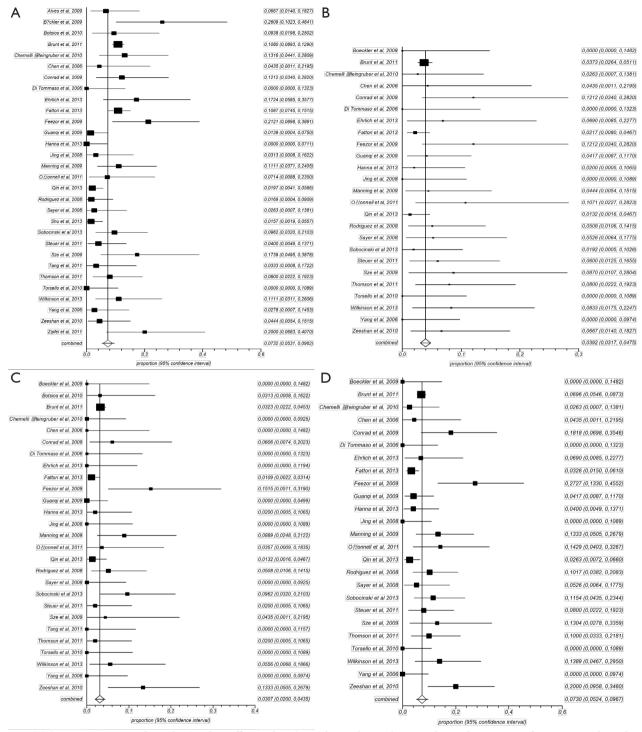
#### Best medical therapy

Outcome data of medical therapy were available for 2,347 patients from 15 studies who underwent conservative medical management for acute type B aortic dissection (*Table 3*). In the vast majority of the papers, indication

for BMT was uncomplicated acute type B dissection. However, a percentage of patients with complications were treated with medical therapy only, either due to the lack of appropriate facilities or due to the presence of comorbidities or morphology that made open surgery or TEVAR not feasible. The 30-day/in-hospital mortality pooled rate was 2.4% (95% CI, 0.9% to 4.6%, *Figure 4A*). The pooled rate for cerebrovascular events was 1% (95% CI, 0.6% to 1.6%), for SCI 0.8% (95% CI, 0.5% to 1.3%) and for overall neurologic complications 2% (95% CI, 0.6% to 4.1%) (*Figure 4B-D*). Survival rates ranged from 86.2% to 100% at 1-year and from 59.0% to 97.2% at 5-years, whereas freedom from aortic events ranged from 34% to 83.9%.

## Endovascular treatment for acute uncomplicated type B dissection

Preliminary results of the European study, Acute Dissection:



**Figure 2** (A) Proportion meta-analysis plot (random effects) of 30-day/in-hospital mortality in endovascular treatment for acute complicated type B dissection. [Pooled proportion, 0.073 (95% CI, 0.053 to 0.096), I<sup>2</sup>, 69.3% (95% CI, 53.4% to 78.1%)]; (B) Forest plot of cerebrovascular events in endovascular treatment for acute complicated type B dissection. Data were available from 25 studies [pooled proportion, 0.039, (95% CI, 0.032 to 0.048), I<sup>2</sup>, 30% (95% CI, 0% to 56.4%)]; (C) Proportion meta-analysis plot (random effects) of spinal cord ischemia in endovascular treatment for acute complicated type B dissection. Data were available from 25 studies [pooled proportion, 0.024), I<sup>2</sup>, 46.3% (95% CI, 6.1% to 64.9%)]; (D) Proportion meta-analysis plot (random effects) of total neurological events in endovascular treatment for acute complicated type B dissection. Data were available from 25 studies [pooled proportion, 0.031, (95% CI, 0.02 to 0.044), I<sup>2</sup>, 46.3% (95% CI, 6.1% to 64.9%)]; (D) Proportion meta-analysis plot (random effects) of total neurological events in endovascular treatment for acute complicated type B dissection. Data were available from 25 studies [pooled proportion, 0.073, (95% CI, 0.052 to 0.097), I<sup>2</sup>, 65.8% (95% CI, 44.1% to 76.6%)].

Table 2 Eligible studies on open surgery for complicated acute type B dissection										
Author	Period of study	N (pts)	CVE (n)	SCI (n)	30-day/in-hospital mortality (n)					
Trimarchi <i>et al.</i> , 2006 (48)	1995-2005	82	8	4	24					
Estrera et al., 2007 (38)	2001-2006	21	ND	ND	4					
Bozinovski et al., 2008 (49)	1989-2004	76	5	5	17					
Shimokawa et al., 2008 (50)	2003-2008	24	ND	1	2					
Zeeshan et al., 2010 (22)	2002-2010	20	0	2	8					
Brunt <i>et al.</i> , 2011 (24)	2005-2008	991	61	25	173					
Murashita et al., 2012 (51)	2003-2010	31	2	2	6					
Minami et al., 2013 (52)	2000-2012	14	4	ND	2					
Wilkinson <i>et al.</i> , 2013 (35)	1995-2012	17	3	1	4					

CVE, cerebrovascular event; SCI, spinal cord ischemia; ND, not determined.

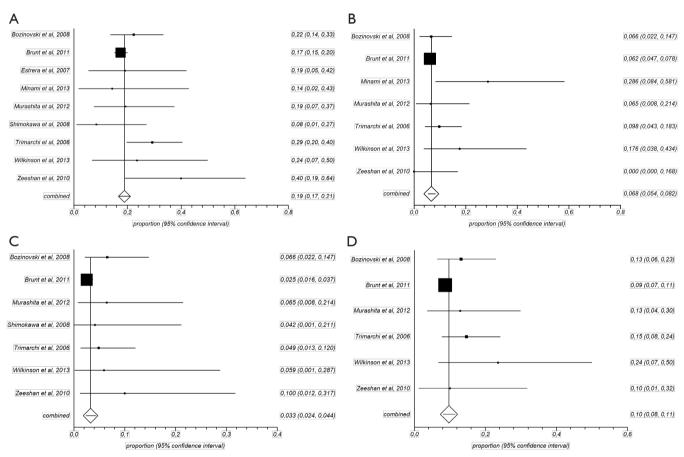


Figure 3 (A) Proportion meta-analysis plot (fixed effects) of 30-day/in-hospital mortality after open surgery for complicated acute type B dissection. [Pooled proportion, 0.19 (95% CI, 0.168 to 0.211), I<sup>2</sup>, 43.6% (95% CI, 0% to 72.4%)]; (B) Proportion meta-analysis plot (fixed effects) of cerebrovascular events after open surgery for complicated acute type B dissection. [Available data in nine studies, pooled proportion, 0.068 (95% CI, 0.054 to 0.082), I<sup>2</sup>, 52% (95% CI, 0% to 77.7%)]; (C) Proportion meta-analysis plot (fixed effects) of spinal cord ischemia after open surgery for complicated acute type B dissection. [Available data in nine studies, pooled proportion, 0.033 (95% CI, 0.024 to 0.044), I<sup>2</sup>, 39.7% (95% CI, 0% to 73.3%)]; (D) Proportion meta-analysis plot (fixed effects) of total neurologic events after open surgery for complicated acute type B dissection. Available data in nine studies, [pooled proportion, 0.098 (95% CI, 0.082 to 0.115), I<sup>2</sup>, 39.4% (95% CI, 0% to 74.8%)].

Table 3 Eligible studies on best medical treatment for uncomplicated acute type B dissection													
Author	Period	Ν	Indication	30-da	ay/in-h	ospital	- FU	Survival rate (%)		Aortic event freedom rate (%)			
Author	of study			CVE	SCI	Mortality		1 year	5 years	1 year	3 years	5 years	10 years
Winnerkvist 2006 (37)	1990- 2001	66	Acute	2	3	0	79	100	82.0	ND	ND	75.0	67.0
Estrera 2007 (38)	2001- 2006	136	Acute not complicated	ND	ND	10	20	ND	75.0	ND	ND	ND	ND
Kitada 2008 (39)	2000- 2006	74	Acute not complicated	ND	ND	0	12	97.0	ND	83.0	ND	ND	ND
Niino 2009 (40)	1996- 2007	210	Acute not complicated	1	1	6	50.5	97.0	89.4	93.0	ND	83.9	76.0
Sakakura 2009 (41)	1996- 2008	215	Acute	ND	1	8	ND	ND	ND	ND	ND	ND	ND
Kitai 2010 (42)	1986- 2008	170	Acute not complicated	ND	1	1	85.2	99.0	85.0	ND	ND	ND	ND
Chemelli- Steingruber 2010 (20)	1996- 2008	50	Acute not complicated	1	ND	3	36	88.0	70.2	88.0	ND	ND	ND
Dick 2010 (43)	2000- 2005	72	Acute not complicated	ND	ND	4	36	ND	79.0	32 need manage		dary surgi	cal
Garbade 2010 (44)	2000- 2008	84	63 not complicated 21 complicated	12	ND	7	36.9	86.2	72.1	22 reinte	erventions	3	
Miyahara 2011 (45)	2000- 2009	160	Acute not complicated	ND	ND	0	33.5	98.7	97.2	92.2	84.2	71.0	ND
Qin <i>et al.</i> 2013 (32)	2004- 2008	41	Acute not com- plicated	0	0	0	40.6	100	59.0	97	63.0	34.0	ND
Fattori 2013 (30)	1995- 2012	853	Acute (315 rupture/ malperfusion)	9	7	74	60	90.2	76.5	82.2	ND	ND	ND
Brunkwall 2013 (46)	2012- 2013	31	Acute not complicated	0	0	0	12	ND	ND	83.9	ND	ND	ND
Nienaber 2013 (5)	2002- 2005	68	Subacute/chronic not complicated	0	0	0	24	97.9	ND	14 TEVA	AR and 4 o	open	
Lu 2013 (47)	1992- 2012	117	Acute not complicated	0	ND	0	58.4	ND	51.2	ND	ND	ND	ND

Table 3 Eligible studies on best medic	I treatment for uncomplicat	ad agute type R dissection
<b>Table 5</b> Eligible studies on Dest medic	I treatment for uncombineat	ed acute type D dissection

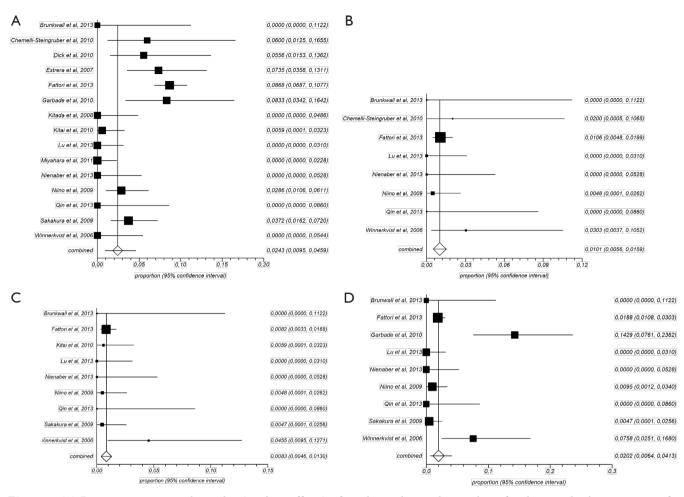
FU, follow-up (months); CVE, cerebrovascular events; SCI, spinal cord ischemia; ND, not determined; TEVAR, treated with endovascular repair.

Stent Graft or Best Medical Treatment (ADSORB) (46,53), which evaluates TEVAR + BMT vs. BMT alone in patients with acute uncomplicated type B aortic dissection showed zero mortality and neurological complication rates in both groups but aortic remodeling after one year was in favor of TEVAR. Another study from China (47) comparing TEVAR + BMT vs. BMT alone in patients with acute uncomplicated type B aortic dissection showed a better long-term survival rate in TEVAR group and confirmed the

favorable aortic remodeling rate (true lumen diameter, false lumen thrombosis).

#### Discussion

Endovascular stent-graft repair of complicated type B acute dissection seems to be associated with favorable short-term and mid-term results. In our analysis, the 30-day/in-hospital mortality was 7.3% in patients treated endovascularly



**Figure 4** (A) Proportion meta-analysis plot (random effects) of 30-day/in-hospital mortality after best medical management for uncomplicated acute type B dissection. [Pooled proportion, 0.024 (95% CI, 0.009 to 0.046), I<sup>2</sup>, 85.9% (95% CI, 78.3% to 90%)]; (B) Proportion meta-analysis plot (fixed effects) of cerebrovascular events after best medical management for uncomplicated acute type B dissection. [Available data in eight studies, pooled proportion, 0.01 (95% CI, 0.006 to 0.016), I<sup>2</sup>, 0% (95% CI, 0% to 56.3%)]; (C) Proportion meta-analysis plot (fixed effects) of spinal cord ischemia after best medical management for uncomplicated acute type B dissection. [Available data in nine studies, pooled proportion, 0.008 (95% CI, 0.005 to 0.013), I<sup>2</sup>, 0% (95% CI, 0% to 54.4%)]; (D) Proportion meta-analysis plot (random effects) of total neurological events after best medical management for uncomplicated acute type B dissection. [Available data in nine studies, pooled proportion, 0.02 (95% CI, 0.006 to 0.013), I<sup>2</sup>, 0% (95% CI, 0% to 54.4%)]; (D) Proportion meta-analysis plot (random effects) of total neurological events after best medical management for uncomplicated acute type B dissection. [Available data in nine studies, pooled proportion, 0.02 (95% CI, 0.006 to 0.041), I<sup>2</sup>, 79.1% (95% CI, 56.2% to 87.5%)].

whereas the pooled rate for 30-day/in-hospital mortality was 19.0% in the operative group. Cerebrovascular events and SCI also occurred more frequently in the operative group. Interestingly, the survival rates for 1- and 5-years were comparable between the two groups. However, the absence of randomized trials comparing endovascular with open repair treatments in complicated type B acute dissection remains a limitation. In addition, it is doubtful whether any diseased anatomy of the dissected aorta can be totally treated by endovascular means. Longer follow-up is warranted to assess the durability of endovascular stentgraft repair for complicated type B acute dissection and potential progression of disease at the downstream aorta.

Uncertainty remains regarding the optimal management strategy for uncomplicated acute type B dissection. The basic medical treatment comprises beta-blockers, diuretics, calcium blockers, angiotensin-converting enzyme inhibitors and alpha-blockers, as well as nitroglycerin. The primary aim of this approach is to obtain a systolic blood pressure between 100 to 120 mmHg and thus reduce the shear stress

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of the aortic wall while maintaining urinary output and visceral perfusion (38). However, BMT is associated with a considerable risk of disease progression to complicated dissection or aneurysmal degeneration of the affected aortic segment, which is the most feared complication, involving about 30-40% of patients (3,54). Long-term outcome of medical therapy has shown a complication rate over 30% and total death rate up to 10% (4).

In our analysis, including predominantly patients with uncomplicated acute type B dissection treated with optimal medical treatment, the pooled 30-day/in-hospital mortality rate was 2.4%. Survival rates ranged from 59.0% to 97.2% at 5-years, whereas freedom from aortic events ranged from 34% to 83.9%, underlining the risk of disease progression to complicated dissection or aneurysmal degeneration of the affected aortic segment in the follow-up and the need for additional surgical interventions. Taking into account the potential risk of disease progression, a crucial question is whether we can expand the indication of endovascular repair in uncomplicated type B aortic dissection and, if so, what are the prognostic factors of early or late complications? The 1-year results of the ADSORB trial showed more frequent false lumen thrombosis and aortic remodeling in those patients treated medically plus TEVAR compared to those managed only medically (53). Favorable aortic remodeling rate was also confirmed by the results of the Investigation of Stent-grafts in Aortic Dissection (INSTEAD-XL) randomized trial which showed that thoracic endovascular aortic repair of uncomplicated type B dissections in addition to medical therapy was associated with improved 5-year aorta-specific survival and delayed disease progression compared to medical therapy alone (5).

The predisposing factors of early and late complications in uncomplicated type B aortic dissection seem to be a critical point that may influence the treatment strategy. Can we identify a high-risk population with uncomplicated type B patients that will benefit from TEVAR? The ideal concept is to perform stent-grafting in the subgroup of patients prone to developing progression of the disease and future complications. A number of studies have suggested several prognostic factors of early or late adverse events such as the patency of the false lumen in the follow-up, an initial aortic diameter  $\geq$ 4 cm with a patent false lumen, an initial false lumen diameter  $\geq 22$  mm in the proximal descending aorta, visceral involvement and recurrent or refractory pain or hypertension (37,42,55-57). Partial false lumen thrombosis, a proximal entry tear size >10 mm and a spiral configuration of the dissection have also been suggested to be associated

with an increased rate of aortic growth. Spiral dissection is associated with a lower incidence of false lumen thrombosis and aorta-related adverse events are more likely to occur in spiral dissection patients. Randomized trials that will focus on these prognostic factors and the optimal timing of intervention are needed.

Our study has the inherent limitations associated with meta-analyses. The considerable heterogeneity amongst identified reports may reflect differing patient characteristics between studies. Hence, these pooled estimates without raw patient data prohibits subset analysis, and our results should be interpreted with caution.

#### Conclusions

Currently, the less invasive method of endovascular repair provides a better 30-day/in-hospital survival for complicated acute type B aortic dissection. Surgical aortic reconstruction, on the other hand, still has a significant role as endovascular repair is not applicable in all of the dissected aortas and there is concern regarding the durability of this technique.

Although the ideal treatment for uncomplicated acute type B aortic dissection is still unclear, the combination of TEVAR with antihypertensive therapy seems to have a more favorable outcome regarding aortic remodeling and aorta-specific survival rate when compared with medical therapy alone. Expansion of TEVAR, however, to treating uncomplicated type B aortic dissection requires further investigation and the proposed factors predicting the outcome of these patients may help in the identification of the appropriate treatment strategy. Until this issue is clarified, patients with uncomplicated acute type B dissection should be treated on a case-by-case basis. There is a need for randomized clinical trials that will focus on the prognostic factors of early and late complications in uncomplicated type B aortic dissection and the timing of intervention.

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#### References

1. Coady MA, Ikonomidis JS, Cheung AT, et al. Surgical management of descending thoracic aortic disease: open and endovascular approaches: a scientific statement

from the American Heart Association. Circulation 2010;121:2780-804.

- 2. Fattori R, Cao P, De Rango P, et al. Interdisciplinary expert consensus document on management of type B aortic dissection. J Am Coll Cardiol 2013;61:1661-78.
- Tsai TT, Trimarchi S, Nienaber CA. Acute aortic dissection: perspectives from the International Registry of Acute Aortic Dissection (IRAD). Eur J Vasc Endovasc Surg 2009;37:149-59.
- Tsai TT, Evangelista A, Nienaber CA, et al. Longterm survival in patients presenting with type A acute aortic dissection: insights from the International Registry of Acute Aortic Dissection (IRAD). Circulation 2006;114:I350-6.
- Nienaber CA, Kische S, Rousseau H, et al. Endovascular repair of type B aortic dissection: long-term results of the randomized investigation of stent grafts in aortic dissection trial. Circ Cardiovasc Interv 2013;6:407-16.
- Stroup DF, Berlin JA, Morton SC, et al. Meta-analysis of observational studies in epidemiology: a proposal for reporting. Meta-analysis Of Observational Studies in Epidemiology (MOOSE) group. JAMA 2000;283:2008-12.
- Di Tommaso L, Monaco M, Mottola M, et al. Major complications following endovascular surgery of descending thoracic aorta. Interact Cardiovasc Thorac Surg 2006;5:705-8.
- 8. Chen S, Yei F, Zhou L, et al. Endovascular stent-grafts treatment in acute aortic dissection (type B): clinical outcomes during early, late, or chronic phases. Catheter Cardiovasc Interv 2006;68:319-25.
- Yang J, Zuo J, Yang L, et al. Endovascular stent-graft treatment of thoracic aortic dissection. Interact Cardiovasc Thorac Surg 2006;5:688-91.
- Jing QM, Han YL, Wang XZ, et al. Endovascular stentgrafts for acute and chronic type B aortic dissection: comparison of clinical outcomes. Chin Med J (Engl) 2008;121:2213-7.
- 11. Sayer D, Bratby M, Brooks M, et al. Aortic morphology following endovascular repair of acute and chronic type B aortic dissection: implications for management. Eur J Vasc Endovasc Surg 2008;36:522-9.
- Rodriguez JA, Olsen DM, Lucas L, et al. Aortic remodeling after endografting of thoracoabdominal aortic dissection. J Vasc Surg 2008;47:1188-94.
- Böckler D, Hyhlik-Dürr A, Hakimi M, et al. Type B aortic dissections: treating the many to benefit the few? J Endovasc Ther 2009;16 Suppl 1:I80-90.
- 14. Alves CM, da Fonseca JH, de Souza JA, et al. Endovascular

treatment of type B aortic dissection: the challenge of late success. Ann Thorac Surg 2009;87:1360-5.

- Conrad MF, Crawford RS, Kwolek CJ, et al. Aortic remodeling after endovascular repair of acute complicated type B aortic dissection. J Vasc Surg 2009;50:510-7.
- Guangqi C, Xiaoxi L, Wei C, et al. Endovascular repair of Stanford type B aortic dissection: early and midterm outcomes of 121 cases. Eur J Vasc Endovasc Surg 2009;38:422-6.
- 17. Feezor RJ, Martin TD, Hess PJ Jr, et al. Early outcomes after endovascular management of acute, complicated type B aortic dissection. J Vasc Surg 2009;49:561-6; discussion 566-7.
- Manning BJ, Dias N, Manno M, et al. Endovascular treatment of acute complicated type B dissection: morphological changes at midterm follow-up. J Endovasc Ther 2009;16:466-74.
- Sze DY, van den Bosch MA, Dake MD, et al. Factors portending endoleak formation after thoracic aortic stent-graft repair of complicated aortic dissection. Circ Cardiovasc Interv 2009;2:105-12.
- 20. Chemelli-Steingruber I, Chemelli A, Strasak A, et al. Endovascular repair or medical treatment of acute type B aortic dissection? A comparison. Eur J Radiol 2010;73:175-80.
- Botsios S, Schuermann K, Maatz W, et al. Complicated acute type B dissections: a single-center experience with endovascular treatment. Thorac Cardiovasc Surg 2010;58:280-4.
- 22. Zeeshan A, Woo EY, Bavaria JE, et al. Thoracic endovascular aortic repair for acute complicated type B aortic dissection: superiority relative to conventional open surgical and medical therapy. J Thorac Cardiovasc Surg 2010;140:S109-15; discussion S142-S146.
- 23. Torsello GB, Torsello GF, Osada N, et al. Midterm results from the TRAVIATA registry: treatment of thoracic aortic disease with the valiant stent graft. J Endovasc Ther 2010;17:137-50.
- Brunt ME, Egorova NN, Moskowitz AJ. Propensity score-matched analysis of open surgical and endovascular repair for type B aortic dissection. Int J Vasc Med 2011;2011:364046.
- 25. Tang JD, Huang JF, Zuo KQ, et al. Emergency endovascular repair of complicated Stanford type B aortic dissections within 24 hours of symptom onset in 30 cases. J Thorac Cardiovasc Surg 2011;141:926-31.
- 26. O'Donnell S, Geotchues A, Beavers F, et al. Endovascular management of acute aortic dissections. J Vasc Surg

2011;54:1283-9.

- Steuer J, Eriksson MO, Nyman R, et al. Early and longterm outcome after thoracic endovascular aortic repair (TEVAR) for acute complicated type B aortic dissection. Eur J Vasc Endovasc Surg 2011;41:318-23.
- Virtue Registry Investigators. The VIRTUE Registry of type B thoracic dissections--study design and early results. Eur J Vasc Endovasc Surg 2011;41:159-66.
- Zipfel B, Czerny M, Funovics M, et al. Endovascular treatment of patients with types A and B thoracic aortic dissection using Relay thoracic stent-grafts: results from the RESTORE Patient Registry. J Endovasc Ther 2011;18:131-43.
- Fattori R, Montgomery D, Lovato L, et al. Survival after endovascular therapy in patients with type B aortic dissection: a report from the International Registry of Acute Aortic Dissection (IRAD). JACC Cardiovasc Interv 2013;6:876-82.
- Ehrlich MP, Rousseau H, Heijmen R, et al. Midterm results after endovascular treatment of acute, complicated type B aortic dissection: the Talent Thoracic Registry. J Thorac Cardiovasc Surg 2013;145:159-65.
- Qin YL, Deng G, Li TX, et al. Treatment of acute type-B aortic dissection: thoracic endovascular aortic repair or medical management alone? JACC Cardiovasc Interv 2013;6:185-91.
- Shu C, Fang K, Luo M, et al. Emergency endovascular stent-grafting for acute type B aortic dissection with symptomatic malperfusion. Int Angiol 2013;32:483-91.
- Hanna JM, Andersen ND, Ganapathi AM, et al. Five-year results for endovascular repair of acute complicated type B aortic dissection. J Vasc Surg 2014;59:96-106.
- Wilkinson DA, Patel HJ, Williams DM, et al. Early open and endovascular thoracic aortic repair for complicated type B aortic dissection. Ann Thorac Surg 2013;96:23-30; discussion 230.
- Sobocinski J, Dias NV, Berger L, et al. Endograft repair of complicated acute type B aortic dissections. Eur J Vasc Endovasc Surg 2013;45:468-74.
- Winnerkvist A, Lockowandt U, Rasmussen E, et al. A prospective study of medically treated acute type B aortic dissection. Eur J Vasc Endovasc Surg 2006;32:349-55.
- Estrera AL, Miller CC, Goodrick J, et al. Update on outcomes of acute type B aortic dissection. Ann Thorac Surg 2007;83:S842-5; discussion S846-50.
- Kitada S, Akutsu K, Tamori Y, et al. Usefulness of fibrinogen/fibrin degradation product to predict poor oneyear outcome of medically treated patients with acute type

B aortic dissection. Am J Cardiol 2008;101:1341-4.

- 40. Niino T, Hata M, Sezai A, et al. Optimal clinical pathway for the patient with type B acute aortic dissection. Circ J 2009;73:264-8.
- 41. Sakakura K, Kubo N, Ako J, et al. Determinants of long-term mortality in patients with type B acute aortic dissection. Am J Hypertens 2009;22:371-7.
- 42. Kitai T, Kaji S, Yamamuro A, et al. Impact of new development of ulcer-like projection on clinical outcomes in patients with type B aortic dissection with closed and thrombosed false lumen. Circulation 2010;122:S74-80.
- 43. Dick F, Hirzel C, Immer FF, et al. Quality of life after acute type B dissection in the era of thoracic endovascular aortic repair. Vasa 2010;39:219-28.
- 44. Garbade J, Jenniches M, Borger MA, et al. Outcome of patients suffering from acute type B aortic dissection: a retrospective single-centre analysis of 135 consecutive patients. Eur J Cardiothorac Surg 2010;38:285-92.
- 45. Miyahara S, Mukohara N, Fukuzumi M, et al. Longterm follow-up of acute type B aortic dissection: ulcer-like projections in thrombosed false lumen play a role in late aortic events. J Thorac Cardiovasc Surg 2011;142:e25-31.
- 46. Brunkwall J. ADSORB was totally underpowered but still shows that AD should be treated. Critical issues in aortic endografting Nuremberg, Germany, 2013.
- Lu Q. Comparison of Endovascular and Medical Therapy in Uncomplicated Stanford Type B Aortic Dissection. Critical issues in aortic endografting Nuremberg, Germany, 2013.
- 48. Trimarchi S, Nienaber CA, Rampoldi V, et al. Role and results of surgery in acute type B aortic dissection: insights from the International Registry of Acute Aortic Dissection (IRAD). Circulation 2006;114:I357-64.
- Bozinovski J, Coselli JS. Outcomes and survival in surgical treatment of descending thoracic aorta with acute dissection. Ann Thorac Surg 2008;85:965-70; discussion 970-1.
- Shimokawa T, Horiuchi K, Ozawa N, et al. Outcome of surgical treatment in patients with acute type B aortic dissection. Ann Thorac Surg 2008;86:103-7.
- 51. Murashita T, Ogino H, Matsuda H, et al. Clinical outcome of emergency surgery for complicated acute type B aortic dissection. Circ J 2012;76:650-4.
- 52. Minami T, Imoto K, Uchida K, et al. Clinical outcomes of emergency surgery for acute type B aortic dissection with rupture. Eur J Cardiothorac Surg 2013;44:360-4; discussion 364-5.
- 53. Brunkwall J, Lammer J, Verhoeven E, et al. ADSORB:

a study on the efficacy of endovascular grafting in uncomplicated acute dissection of the descending aorta. Eur J Vasc Endovasc Surg 2012;44:31-6.

- 54. Nienaber CA, Rousseau H, Eggebrecht H, et al. Randomized comparison of strategies for type B aortic dissection: the INvestigation of STEnt Grafts in Aortic Dissection (INSTEAD) trial. Circulation 2009;120:2519-28.
- 55. Marui A, Mochizuki T, Koyama T, et al. Degree of fusiform dilatation of the proximal descending aorta in

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type B acute aortic dissection can predict late aortic events. J Thorac Cardiovasc Surg 2007;134:1163-70.

- 56. Song JM, Kim SD, Kim JH, et al. Long-term predictors of descending aorta aneurysmal change in patients with aortic dissection. J Am Coll Cardiol 2007;50:799-804.
- 57. Trimarchi S, Eagle KA, Nienaber CA, et al. Importance of refractory pain and hypertension in acute type B aortic dissection: insights from the International Registry of Acute Aortic Dissection (IRAD). Circulation 2010;122:1283-9.

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