## Drug-eluting stents versus coronary artery bypass grafting for the treatment of coronary artery disease: A meta-analysis of randomized and nonrandomized studies

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**Background:** We performed the present systematic review and meta-analysis of randomized and nonrandomized comparative studies in an attempt to compare the safety and efficacy of drug-eluting stents with coronary artery bypass grafting for patients with coronary artery disease.

**Methods:** Twenty-five eligible comparative studies (1 randomized and 24 nonrandomized) were assessed. Two reviewers independently appraised each study. Meta-analysis was performed by combining the results of reported incidence of morbidity, mortality, and repeat revascularization. The relative risk was used as a summary statistic.

**Results:** In these 25 studies 34,278 patients were compared, of whom 18,538 received drug-eluting stents and 15,740 underwent coronary artery bypass grafting. It must be acknowledged that this comparison represented a selected group of patients who received drug-eluting stents or underwent coronary artery bypass grafting. The accumulative incidences of all-cause mortality at 12 months (4.5% vs 4.0%, P = .92) and 24 months (6.2% vs 8.4%, P = .27) and 30-day myocardial infarction (1.4% vs 2.0%, P = .60) were similar, respectively, between the drug-eluting stent and coronary artery bypass grafting groups. Drug-eluting stents were associated with lower rates of all-cause mortality at 30 days (0.9% vs 2.3%, P < .001), stroke (0.4% vs 1.7%, P < .001), and 30-day major adverse cardiac and cerebrovascular events (3.6% vs 5.5%, P < .04). However, the coronary artery bypass grafting group had a lower incidence of postprocedural myocardial infarction (5.5% vs 4.7%, P = .03), repeat revascularization (22.2% vs 4.1%, P < .001), and 12-month major adverse cardiac and cerebrovascular events (16.7% vs 10.5%, P < .001). Subgroup analysis of patients with multivessel coronary artery disease showed similar results.

**Conclusions:** Drug-eluting stents are associated with less periprocedural risks but a higher incidence of postprocedural myocardial infarction, repeat revascularization, and 12-month major adverse cardiac and cerebrovascular events compared with coronary artery bypass grafting. (J Thorac Cardiovasc Surg 2011;141:1134-44)



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Coronary artery bypass grafting (CABG) and percutaneous coronary intervention (PCI) are alternative revascularization techniques for symptomatic patients with coronary artery disease.<sup>1</sup> Increased experience with PCI and improved

Disclosures: A uthors have nothing to disclose with regard to commercial support.

0022-5223/\$36.00

doi:10.1016/j.jtcvs.2010.07.001

technological advent have expanded its use in patients with severe coronary artery disease and complex anatomic lesions. A meta-analysis by Mercado and colleagues<sup>2</sup> comparing randomized controlled trials (RCTs) of PCI with bare metal stents (BMSs) versus CABG demonstrated similar degrees of protection against death, myocardial infarction, and stroke for patients with multivessel coronary artery disease at 1 year after the initial procedure. However, repeat revascularization procedures remained more likely after use of BMSs. A subsequent meta-analysis by Daemen and coworkers<sup>3</sup> comparing RCTs on the long-term outcomes between PCI with BMSs and CABG (the Stent of Surgery trial, the Arterial Revascularization Therapies Study, Edstudio Randomizado Argentino de Angioplastia vs Cirugia II, and the Medicine, Angioplasty or Surgery Study II) indicated that BMSs were associated with a long-term safety profile similar to that of CABG but also reinforced higher revascularization and major adverse cardiac and cerebrovascular events (MACCEs) in the BMS group.<sup>3</sup>

Drug-eluting stents (DES) demonstrate similar rates of death and myocardial infarction but reduced rates of repeat

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Received for publication Nov 26, 2009; revisions received June 22, 2010; accepted for publication July 6, 2010; available ahead of print Dec 20, 2010.

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| Abbreviations and Acronyms |                                      |  |  |  |  |  |  |  |
|----------------------------|--------------------------------------|--|--|--|--|--|--|--|
| BMS                        | = bare metal stent                   |  |  |  |  |  |  |  |
| CABG                       | = coronary artery bypass grafting    |  |  |  |  |  |  |  |
| CI                         | = confidence interval                |  |  |  |  |  |  |  |
| DES                        | = drug-eluting stent                 |  |  |  |  |  |  |  |
| MACCE                      | = major adverse cardiac and          |  |  |  |  |  |  |  |
|                            | cerebrovascular event                |  |  |  |  |  |  |  |
| PCI                        | = percutaneous coronary intervention |  |  |  |  |  |  |  |
| RCT                        | = randomized controlled trial        |  |  |  |  |  |  |  |
| RR                         | = relative risk                      |  |  |  |  |  |  |  |
| SYNTAX                     | = Synergy Between PCI with Taxus     |  |  |  |  |  |  |  |
|                            | and Cardiac Surgery                  |  |  |  |  |  |  |  |

vessel or left main coronary artery disease.<sup>6</sup> This RCT demonstrated that at 12 months the rates of death and myocardial infarction were similar between the 2 groups, but DESs were associated with a significantly higher rate of MACCEs (17.8% vs 12.4%, P = .002) and lower rate of stroke (0.6% vs 2.2%, P = .003). As is generally true with RCTs, the study population is predefined and hence subject to trial design bias. We performed the present systematic review and meta-analysis of the randomized and nonrandomized comparative studies in an attempt to assess the safety and efficacy of DESs versus CABG with the current clinical evidence.

## METHODS

## Search Strategy

revascularization compared with BMSs,<sup>4</sup> thus increasing the percentage of patients with multivessel disease treated with PCI. However, recent data suggested a higher rate of thrombotic occlusion with DESs than BMSs.<sup>5</sup> The recent Synergy Between PCI with Taxus and Cardiac Surgery (SYNTAX) trial compared DESs versus CABG in patients with triple-

Electronic searches were performed in 6 databases from their inception to September 2009: Medline, Embase, Pubmed, Cochrane Central Register of Controlled Trials, Cochrane Database of Systematic Reviews, and Database of Abstracts of Review of Effectiveness. To achieve the maximum sensitivity of the search strategy and identify all trials comparing DESs and CABG, we used appropriate free text and thesaurus terms, including "percutaneous coronary intervention," "coronary angioplasty," "coronary artery stenting," "drug-eluting stent," and "coronary artery bypass

|  |              |                     |              | No. of patients |      | Age of cohort<br>(mean ± SD) |           | Ejection fraction<br>(mean ± SD) |             |
|--|--------------|---------------------|--------------|-----------------|------|------------------------------|-----------|----------------------------------|-------------|
| Reference                                    | Study design | Study population    | Study period | DES             | CABG | DES                          | CABG      | DES                              | CABG        |
| Ben-Gal and coworkers9                       | OC           | DM                  | 2002-2005    | 86              | 86   | NA                           | NA        | NA                               | NA          |
| Ben-Gal and coworkers <sup>10</sup>          | OC           | LAD                 | 2002-2003    | 83              | 83   | NA                           | NA        | NA                               | NA          |
| Briguori and coworkers <sup>11</sup>         | OC           | MVD, LAD, DM, OPCAB | 2002-2004    | 69              | 149  | $63\pm9$                     | $66\pm9$  | $54\pm12$                        | $53\pm9$    |
| Cheng and coworkers <sup>12</sup>            | OC           | LMCAD               | 2000-2007    | 94              | 216  | $68\pm10$                    | $67\pm9$  | $56\pm17$                        | $56\pm20$   |
| Chieffo and coworkers <sup>13</sup>          | OC           | LMCAD               | 2002-2004    | 107             | 142  | $64\pm10$                    | $68\pm10$ | $52\pm10$                        | $52\pm11$   |
| Domínguez-Franco and coworkers <sup>14</sup> | OC           | MVD, LAD, DM        | 2000-2004    | 128             | 142  | 68                           | 65        | $52\pm13$                        | $54 \pm 14$ |
| Gioia and coworkers <sup>15</sup>            | OC           | LV dysfunction      | 2002-2005    | 128             | 92   | $69\pm10$                    | $68\pm10$ | $28\pm 6$                        | $27\pm8$    |
| Hannan and coworkers <sup>16</sup>           | OC           | MVD                 | 2003-2005    | 9963            | 7437 | $65\pm12$                    | $66\pm11$ | NA                               | NA          |
| Hong and coworkers <sup>17</sup>             | OC           | LAD                 | 2003         | 119             | 70   | $61\pm10$                    | $61\pm10$ | $53\pm9$                         | $52\pm9$    |
| Javaid and coworkers <sup>18</sup>           | OC           | MVD                 | NA           | 979             | 701  | $66\pm11$                    | $65\pm11$ | NA                               | NA          |
| Kukreja and coworkers <sup>19</sup>          | OC           | MVD, LAD            | 1997-2003    | 289             | 206  | $63\pm10$                    | $62\pm9$  | $59\pm12$                        | $61\pm13$   |
| White and coworkers <sup>20</sup>            | OC           | LMCAD               | 2003-2005    | 50              | 123  | $72\pm15$                    | $70\pm10$ | $51\pm15$                        | $52\pm10$   |
| Li and coworkers <sup>21</sup>               | OC           | MVD                 | 2004-2005    | 1834            | 1886 | $58\pm10$                    | $61\pm9$  | NA                               | NA          |
| Mäkikallio and coworkers <sup>22</sup>       | OC           | LMCAD               | 2005-2007    | 49              | 238  | $72\pm10$                    | $70\pm9$  | $55\pm12$                        | $54\pm11$   |
| Moshkovitz and coworkers <sup>23</sup>       | OC           | LAD, OPCAB          | 2002-2003    | 116             | 116  | NA                           | NA        | NA                               | NA          |
| Palmerini and coworkers <sup>24</sup>        | OC           | LMCAD               | 2003-2006    | 98              | 161  | 81*                          | 78*       | 50*                              | 53*         |
| Park and coworkers <sup>25</sup>             | OC           | MVD                 | 2003-2005    | 1547            | 1495 | $62\pm10$                    | $62\pm9$  | $59\pm9$                         | $56\pm11$   |
| Sanmartín and coworkers <sup>26</sup>        | OC           | LMCAD               | 2000-2005    | 96              | 245  | $66\pm13$                    | $66\pm10$ | NA                               | NA          |
| Serruys and coworkers <sup>6</sup>           | RCT          | MVD or LMCAD        | 2005-2007    | 903             | 897  | $65\pm10$                    | $65\pm10$ | NA                               | NA          |
| Tarantini and coworkers <sup>27</sup>        | OC           | MVD, DM             | 2004-2005    | 93              | 127  | $65\pm9$                     | $67\pm7$  | $62\pm14$                        | $62\pm14$   |
| Toutouzas and coworkers <sup>28</sup>        | OC           | LAD, DM             | 2001-2006    | 39              | 38   | $59\pm13$                    | $61\pm10$ | $48\pm7$                         | $49\pm9$    |
| van Domburg and coworkers <sup>29</sup>      | OC           | MVD or LMCAD        | 2002         | 798             | 275  | 62 ± 11                      | 64 ± 11   | NA                               | NA          |
| Yang <sup>30</sup>                           | OC           | MVD                 | 2003-2004    | 235             | 231  | $65\pm10$                    | $65\pm10$ | $51\pm9$                         | $50\pm11$   |
| Yang and coworkers <sup>31</sup>             | OC           | MVD                 | 2003-2005    | 441             | 390  | $63\pm10$                    | $63\pm 8$ | $58\pm12$                        | $53\pm14$   |
| Yi and coworkers <sup>32</sup>               | OC           | MVD, OPCAB          | 2003-2005    | 194             | 194  | $63\pm10$                    | $62\pm9$  | NA                               | NA          |

*SD*, Standard deviation; *DES*, drug-eluting stent; *CABG*, coronary artery bypass grafting; *OC*, observational cohort; *DM*, diabetes mellitus; *NA*, not applicable; *LAD*, left anterior descending coronary artery; *MVD*, multivessel disease; *OPCAB*, off-pump coronary artery bypass; *LMCAD*, left main coronary artery disease; *LV*, left ventricle; *RCT*, randomized controlled trial \*Median

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