Determinants of variations in initial treatment strategies for stable ischemic heart disease

Maria C. Bennell MPH, Feng Qiu MSc, Kori J. Kingsbury MSN, Peter C. Austin PhD, Harindra C. Wijeysundera MD

Abstract

Background: The ratio of revascularization to medical therapy (referred to herein as the revascularization ratio) for the initial treatment of stable ischemic heart disease varies considerably across hospitals. We conducted a comprehensive study to identify patient, physician and hospital factors associated with variations in the revascularization ratio across 18 cardiac centres in the province of Ontario. We also explored whether clinical outcomes differed between hospitals with high, medium and low ratios.

Methods: We identified all patients in Ontario who had stable ischemic heart disease documented by index angiography performed between Oct. 1, 2008, and Sept. 30, 2011, at any of the 18 cardiac centres in the province. We classified patients by initial treatment strategy (medical therapy or revascularization). Hospitals were classified into equal tertiles based on their revascularization ratio. The primary outcome was all-cause mortality. Patient follow-up was until Dec. 31, 2012. Hierarchical logistic regression models identified predictors of revascularization. Multivariable Cox proportional hazards models, with a time-varying covariate for actual treatment received, were used to evaluate the impact of the revascularization ratio on clinical outcomes.

Results: Variation in revascularization ratios was twofold across the hospitals. Patient factors accounted for 67.4% of the variation in revascularization ratios. Physician and hospital factors were not significantly associated with the variation. Significant patient-level predictors of revascularization were history of smoking, multivessel disease, high-risk findings on noninvasive stress testing and more severe symptoms of angina (v. no symptoms). Treatment at hospitals with a high revascularization ratio was associated with increased mortality compared with treatment at hospitals with a low ratio (hazard ratio 1.12, 95% confidence interval 1.03–1.21).

Interpretation: Most of the variation in revascularization ratios across hospitals was warranted, in that it was driven by patient factors. Nonetheless, the variation was associated with potentially important differences in mortality.

Stable ischemic heart disease is a common manifestation of cardiovascular disease, the leading cause of death in the world.1,2 The treatment strategies for stable ischemic heart disease include medical therapy alone or in combination with revascularization by percutaneous coronary intervention (PCI) or coronary artery bypass grafting (CABG).

A tremendous amount of research has examined the best initial treatment strategy for stable ischemic heart disease.3–5 Randomized controlled trials have not shown a difference in major adverse events between optimal medical therapy and revascularization.6 Some argue that revascularization should be reserved only for symptom relief.7,8 Criteria for the appropriate use of revascularization have been developed to aid in clinical decision-making; however, a substantial proportion of revascularization procedures for stable ischemic heart disease are performed under clinical circumstances deemed as “uncertain.”9,10 Reflecting this uncertainty, there is wide regional variation in the rate of coronary revascularization,11–13 which suggests different thresholds for invasive therapy for stable ischemic heart disease.

Studies have predominantly examined the determinants of variations in the type of revascularization modality used.13,14 There is a paucity of data exploring the determinants of variations in the earlier decision to treat with medical therapy alone or with revascularization. A study published nearly a decade ago did not examine outcomes.7 Accordingly, our primary research objective was to determine whether the variations in initial...
Table 1 (part 1 of 2): Baseline patient, physician and hospital characteristics by initial treatment strategy for stable ischemic heart disease

| Characteristic                          | Initial treatment strategy; % of patients* |
|----------------------------------------|---------------------------------------------|
|                                        | Total | Medical therapy | Revascularization | p value†          |
|                                        | n = 39 131 | n = 15 139 | n = 23 992         |                  |
| **Patient**                            |       |               |                  |                  |
| Age, yr, mean ± SD                     | 66.0 ± 10.3 | 67.0 ± 10.2 | 65.3 ± 10.3       | < 0.001          |
| Male sex                               | 75.3    | 74.3         | 75.9             | < 0.001          |
| Rural residence                        | 14.5    | 14.4         | 14.5             | 0.7              |
| Income quintile                        |        |               |                  | < 0.001          |
| 1 (lowest)                             | 18.6    | 19.6         | 17.9             |                  |
| 2                                      | 20.4    | 20.4         | 20.4             |                  |
| 3                                      | 20.2    | 20.3         | 20.2             |                  |
| 4                                      | 20.7    | 20.2         | 21.0             |                  |
| 5 (highest)                            | 20.1    | 19.4         | 20.5             |                  |
| **Medical comorbidity**                |       |               |                  |                  |
| Peripheral vascular disease            | 9.4     | 11.5         | 8.1              | < 0.001          |
| Prior MI                               | 28.1    | 35.8         | 23.3             | < 0.001          |
| COPD                                   | 6.9     | 8.6          | 5.8              | < 0.001          |
| Charlson comorbidity score, mean ± SD  | 0.9 ± 1.3 | 1.1 ± 1.5 | 0.8 ± 1.2        | < 0.001          |
| **Cardiac risk factors**               |       |               |                  |                  |
| Diabetes                               | 44.0    | 48.0         | 41.4             | < 0.001          |
| Hypertension                           | 86.7    | 89.5         | 84.9             | < 0.001          |
| Hyperlipidemia                         | 80.8    | 83.0         | 79.5             | < 0.001          |
| History of smoking                     | 31.6    | 33.2         | 30.5             | < 0.001          |
| **Cardiac status/testing**             |       |               |                  |                  |
| Native stenosis‡                       |        |               |                  |                  |
| Left main coronary artery              | 13.0    | 10.8         | 14.4             | < 0.001          |
| Proximal left anterior descending coronary artery | 33.2 | 28.7 | 36.0 | < 0.001 |
| Mid or distal left anterior descending coronary artery | 49.6 | 47.5 | 51.0 | < 0.001 |
| Circumflex artery                      | 51.5    | 50.9         | 51.9             | 0.07             |
| Right coronary artery                  | 60.8    | 60.5         | 60.9             | 0.4              |
| Prior CABG                             |        |               |                  | < 0.001          |
| Yes                                    | 18.0    | 30.1         | 10.4             |                  |
| No                                     | 81.9    | 69.9         | 89.5             |                  |
| Unknown                                | 0.1     | 0.1          | 0.1              |                  |
| Vessel disease                         |        |               |                  | < 0.001          |
| 1 vessel                               | 38.9    | 42.0         | 36.9             |                  |
| 2 vessels                              | 25.8    | 24.0         | 27.0             |                  |
| 3 vessels                              | 34.5    | 32.5         | 35.8             |                  |
| Missing data                           | 0.8     | 1.5          | 0.3              |                  |
| Left ventricular function              |        |               |                  | < 0.001          |
| ≤ 34%                                  | 5.4     | 8.4          | 3.5              |                  |
| 35%–49%                                | 12.9    | 16.0         | 11.0             |                  |
| ≥ 50%                                  | 48.6    | 46.5         | 50.0             |                  |
| Not done                               | 33.0    | 29.1         | 35.5             |                  |
Research

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Organic treatment strategies for stable ischemic heart disease are warranted. We conducted a comprehensive population-based study to identify patient, physician and hospital factors associated with variations in treatment strategies within 90 days after angiography. We also explored whether clinical outcomes differed between hospitals with high, medium and low ratios of revascularization to medical therapy (hereafter referred to as the revascularization ratio).

Methods

Study design and setting
We conducted an observational cohort study. Our primary data source was the Cardiac Care

Table 1 (part 2 of 2): Baseline patient, physician and hospital characteristics by initial treatment strategy for stable ischemic heart disease

| Characteristic | Total n = 39 131 | Medical therapy n = 15 139 | Revascularization n = 23 992 | p value† |
|----------------|------------------|-----------------------------|-----------------------------|----------|
| Exercise ECG risk |                  |                             |                             | < 0.001 |
| High risk       | 27.7             | 21.3                        | 31.7                        |          |
| Low risk        | 22.4             | 22.9                        | 22.2                        |          |
| Uninterpretable | 4.9              | 4.8                         | 4.9                         |          |
| Not done        | 45.0             | 51.1                        | 41.2                        |          |
| Functional imaging risk |             |                             |                             | < 0.001 |
| High risk       | 32.0             | 31.2                        | 32.6                        |          |
| Low risk        | 23.0             | 25.1                        | 21.7                        |          |
| Not done        | 44.9             | 43.7                        | 45.7                        |          |
| CCS class       |                  |                             |                             | < 0.001 |
| 0               | 16.5             | 22.0                        | 13.0                        |          |
| 1               | 14.0             | 15.9                        | 12.8                        |          |
| 2               | 38.3             | 35.5                        | 40.0                        |          |
| 3               | 28.5             | 24.2                        | 31.1                        |          |
| 4               | 2.8              | 2.4                         | 3.0                         |          |
| Physician§      |                  |                             |                             |          |
| Age, yr, mean ± SD | 48.3 ± 9.1      | 48.8 ± 9.1                  | 47.9 ± 9.0                  | < 0.001 |
| Sex             |                  |                             |                             |          |
| Female          | 3.7              | 3.9                         | 3.6                         |          |
| Male            | 94.7             | 94.5                        | 94.9                        |          |
| Unknown         | 1.5              | 1.6                         | 1.5                         |          |
| Years since graduation, mean ± SD | 23.4 ± 9.3 | 23.9 ± 9.3 | 23.1 ± 9.3 | < 0.001 |
| Total consultations billed per year, mean ± SD | 856.5 ± 424.7 | 863.0 ± 424.5 | 852.4 ± 424.8 | 0.02 |
| Total visits billed per year, mean ± SD | 2613.9 ± 1730.7 | 2707.5 ± 1773.6 | 2555.0 ± 1700.5 | < 0.001 |
| Hospital        |                  |                             |                             |          |
| Annual catheterization volume, mean ± SD | 4092.3 ± 1696.4 | 4031.3 ± 1684.3 | 4130.9 ± 1702.9 | < 0.001 |
| Hospital type   |                  |                             |                             | < 0.001 |
| Catheterization only | 13.2        | 13.6                        | 13.0                        |          |
| PCI and catheterization only | 6.1          | 7.2                         | 5.4                         |          |
| CABG, PCI and catheterization | 80.8         | 79.3                        | 81.7                        |          |

Note: CABG = coronary artery bypass grafting, CCS = Canadian Cardiovascular Society, COPD = chronic obstructive pulmonary disease, ECG = electrocardiogram, MI = myocardial infarction, PCI = percutaneous coronary intervention, SD = standard deviation.

*Unless stated otherwise.
†We assessed differences in baseline characteristics between initial treatment strategy and hospital revascularization ratio strata using the χ² test for categorical variables and analysis of variance for continuous variables.
‡Significant stenosis ≥ 70% (≥ 50% in left main coronary artery).
§Procedural physician.
Network of Ontario database. The database receives clinical data from the 18 cardiac centres in the province on all patients who undergo cardiac angiography, PCI or CABG. The database’s accuracy has been previously validated by means of retrospective chart review and comparisons with other databases. We linked the data to those from population-level administrative databases with the use of unique, encoded identifiers (the list of administrative databases is available in Appendix 1, www.cmaj.ca/lookup/suppl/doi:10.1503/cmaj.141372/-/DC1).

The Research Ethics Board at the Sunnybrook Health Sciences Centre, University of Toronto, approved the study design.

Study population
We included Ontario residents 20 years of age and over who had stable ischemic heart disease documented by index angiography between Oct. 1, 2008, and Sept. 30, 2011, at any of the 18 cardiac centres. For patients who underwent multiple angiograms within the study period, we included only the first angiogram. To use a conservative definition, we excluded patients with normal coronary arteries, mild coronary artery disease (stenosis < 70%, or < 50% in the left main artery) or a recent myocardial infarction (≤ 90 d before the index angiogram). Patients who underwent PCI or CABG within 90 days after their index angiogram were included in the revascularization group; the remaining patients were included in the medical therapy group. We chose 90 days as the cut-off because it is consistent with current procedural wait times.

Determination of revascularization ratios
We calculated the revascularization ratio for each hospital based on the number of patients who had an index angiogram at that hospital and were subsequently allocated to either revascularization or medical therapy. We then classified the participating hospitals into 3 categories by ratio (high, medium or low) such that there were equal numbers of hospitals in each category.

Outcome measures
The primary outcome was all-cause mortality, and the secondary outcome was readmission with nonfatal myocardial infarction. Maximum follow-up was until Dec. 31, 2012.

Statistical analysis
We found that 1.5% of patients had missing data on details of the procedural physician, and 0.1% had missing data on whether they had a prior CABG. We excluded these patients from the subsequent regression modelling. We used a 3-level hierarchical logistic regression model to identify drivers of revascularization. To account for clustering of patients by procedural physician and by procedural hospital, we included hospital-specific and physician-specific random effects, with the procedural physician nested within the procedural hospital. We developed 5 models: an unconditional/null model that contained the physician- and hospital-specific random effects only; a model adjusted for significant patient characteristics; a model adjusted for significant patient and physician characteristics; a model adjusted for significant patient and hospital characteristics; and finally a model adjusted for all variables.

The variance of the distribution of the hospital-specific random effect is a measure of the

| Table 2: Distribution of participating hospitals by hospital revascularization ratio* |
|---------------------------------|-------------------------------|
| Ratio category; hospital        | Revascularization ratio       |
| Low ratio                       |
| Hôtel-Dieu Grace Hospital (Windsor) | 1.09                         |
| St. Mary's General Hospital (Kitchener) | 1.18                         |
| Brampton Civic Hospital         | 1.26                         |
| Southlake Regional Health Centre (Newmarket) | 1.31                         |
| Thunder Bay Regional Health Sciences Centre | 1.34                        |
| University Health Network (Toronto) | 1.41                        |
| Medium ratio                    |
| Peterborough Regional Health Centre | 1.44                        |
| St. Michael's Hospital (Toronto) | 1.45                         |
| Rouge Valley Health System (Toronto) | 1.46                        |
| Sudbury Regional Hospital (now Health Sciences North) | 1.58                        |
| London Health Sciences Centre   | 1.60                         |
| Trillium Health Centre (Mississauga) | 1.63                        |
| High ratio                      |
| Sault Area Hospital             | 1.69                         |
| Hamilton Health Sciences        | 1.76                         |
| Toronto East General Hospital   | 1.86                         |
| Sunnybrook Health Sciences Centre (Toronto) | 1.96                        |
| Kingston General Hospital       | 2.13                         |
| University of Ottawa Heart Institute | 2.31                        |

*Ratio of revascularization to medical therapy.
between-hospital variation in the choice of initial treatment strategy. We examined the proportional change in the variance of the random effect between models 1 and 2, 2 and 3, 2 and 4, and 3 and 5, and identified important predictors of revascularization using model 5.

We compared unadjusted clinical outcomes across the 3 categories of hospital revascularization ratio using Kaplan–Meier curves. We developed adjusted Cox-proportional hazards models using a time-varying covariate for treatment received to address the potential for immortal time bias. The entire cohort was included in the medical therapy category until the time of revascularization (up to a maximum of 90 d), at which point they were moved to the PCI or CABG group based on the first revascularization modality received. We used robust “sandwich-type” variance estimators to account for potential homogeneity in outcomes for clusters of patients treated at the same hospital. We adjusted for patient, physician and hospital characteristics and evaluated the association between the revascularization ratio and clinical outcomes. We compared hospitals with high, medium and low revascularization ratios within each treatment strategy by means of an interaction term.

We used SAS version 9.3 (SAS Institute Inc.) for the statistical analyses.

Results

We identified 39 131 patients with stable ischemic heart disease who met our inclusion criteria; 15 139 of them received medical therapy, and 23 992 underwent revascularization. The baseline characteristics of the patients are reported in Table 1. There were numerous differences between patients in the medical therapy group and those in the revascularization group, with the latter on average being younger and having fewer comorbidities, more diseased vessels and more severe angina symptoms. (The baseline characteristics of patients in the revascularization group by type of procedure are available in Appendix 2, at www.cmaj.ca/lookup/suppl/doi:10.1503/cmaj.141372/-/DC1.)

The mean overall revascularization ratio was 1.58. We observed a twofold variation in the revascularization ratios across the 18 hospitals, from 1.09 to 2.31 (Table 2). In general, patients were relatively similar, with no clinically meaningful differences across the 3 revascularization ratio groups (Appendix 3, available at www.cmaj.ca/lookup/suppl/doi:10.1503/cmaj.141372/-/DC1).

Patient factors significantly associated with an increased likelihood of revascularization instead of medical therapy were history of smoking (odds ratio [OR] 1.07, 95% confidence interval [CI] 1.01–1.12), multivessel disease (2 vessels: OR 1.45, 95% CI 1.34–1.57; 3 vessels: OR 1.66, 95% CI 1.47–1.88), high risk findings on noninvasive stress testing (OR 1.27, 95% CI 1.18–1.36) and more severe symptoms of angina (OR 1.36–2.38 for Canadian Cardiovascular Society [CCS] classes 1–4 relative to CCS class 0) (Table 3). Coronary anatomy was also a strong predictor of revascularization, in particular stenoses in the left main artery (OR 1.51, 95% CI 1.37–1.67) and the proximal left anterior descending artery (OR 1.69, 95% CI 1.59–1.79). In contrast, older patients and those with more medical comorbidities and cardiac risk factors were less likely to undergo revascularization. In particular, prior CABG (OR...
0.19, 95% CI 0.18–0.20) and poor left ventricular function (OR 0.50, 95% CI 0.45–0.55) were strong predictors of medical therapy (Table 3).

Of the physician factors, age was the only significant factor: patients who underwent an angiogram by an older physician were less likely to undergo revascularization (OR 0.97, 95% CI 0.94–0.99, per year increase) (Table 3). Hospital factors did not significantly predict likelihood of revascularization.

The addition of patient variables reduced the variance of the hospital random effect from 0.0224 in the unconditional model to 0.0073, for a reduction of 67.4%. Thus, the between-hospital variation in revascularization strategy for a reference patient (one whose patient covariates were all set to zero) was less than the between-hospital variation in crude revascularization ratio. Further addition of physician and hospital variables did not reduce the between-hospital variation in initial treatment strategy (variance estimate 0.0093 after addition of physician variables, 0.0094 after addition of hospital variables, and 0.0116 after addition of physician and hospital variables).

Over a median follow-up of 2.5 years, there was no difference in unadjusted all-cause mortality ($p = 0.3$) or readmission with nonfatal myocardial infarction ($p = 0.09$) between the 3 hospital revascularization ratio categories (Figure 1). In the risk-adjusted models, patients treated at a hospital in the high revascularization ratio category were 12% more likely to die than those treated at a centre in the low ratio category (hazard ratio [HR] 1.12, 95% CI 1.03–1.21) (Table 4). There was no statis-

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### Table 3: Factors associated with the likelihood of revascularization instead of medical therapy among patients with stable ischemic heart disease

| Factor                                      | OR (95% CI)          | Factor                                      | OR (95% CI)          |
|---------------------------------------------|----------------------|---------------------------------------------|----------------------|
| **Patient**                                 |                      | **Left ventricular function**               |                      |
| Age (per additional 10 yr)                  | 0.89 (0.87–0.91)     | ≤34%                                        | 0.50 (0.45–0.55)     |
| Male (v. female)                            | 1.05 (0.99–1.10)     | 35%–49%                                     | 0.83 (0.77–0.89)     |
| Rural residence (v. urban)                  | 0.96 (0.90–1.03)     | ≥50%                                        | 1.00 (ref)           |
| (per additional point)                      |                      | Not done                                    | 1.08 (1.01–1.14)     |
| **Medical comorbidities**                   |                      | **Exercise ECG risk**                       |                      |
| Peripheral vascular disease (v. no disease)  | 0.85 (0.79–0.92)     | Low risk                                    | 1.00 (ref)           |
| Prior MI (v. no prior MI)                   | 0.79 (0.74–0.83)     | High risk                                   | 1.27 (1.18–1.36)     |
| COPD (v. no COPD)                          | 0.84 (0.76–0.91)     | Uninterpretable                             | 1.09 (0.97–1.22)     |
| Charlson comorbidity score                  | 0.92 (0.91–0.94)     | Not done                                    | 0.94 (0.88–1.00)     |
| **Cardiac risk factors**                    |                      | **Functional imaging risk**                 |                      |
| Diabetes (v. no diabetes)                   | 0.94 (0.89–0.99)     | Low risk                                    | 1.00 (ref)           |
| Hypertension (v. no hypertension)           | 0.89 (0.83–0.96)     | High risk                                   | 1.07 (1.00–1.14)     |
| Hyperlipidemia (v. no hyperlipidemia)       | 0.94 (0.89–1.00)     | Uninterpretable                             | 1.12 (1.05–1.20)     |
| History of smoking (v. never smoked)        | 1.07 (1.01–1.12)     | Not done                                    | 1.12 (1.05–1.20)     |
| **Cardiac status testing**                  |                      | **CCS class**                               |                      |
| Native stenosis* (v. nonsignificant stenosis)|                      | 0                                           | 1.00 (ref)           |
| Left main coronary artery                   | 1.51 (1.37–1.67)     | 1                                           | 1.36 (1.25–1.47)     |
| Proximal left anterior descending coronary artery | 1.69 (1.59–1.79) | 2                                           | 1.81 (1.69–1.93)     |
| Mid or distal left anterior descending coronary artery | 1.18 (1.11–1.25) | 3                                           | 2.31 (2.14–2.48)     |
| Circumflex artery                          | 1.03 (0.96–1.10)     | 4                                           | 2.38 (2.04–2.78)     |
| Right coronary artery                      | 1.09 (1.02–1.17)     | **Physician†**                              |                      |
| Prior CABG (v. no prior CABG)               | 0.19 (0.18–0.20)     | Age, yr (per additional yr)                | 0.97 (0.94–0.99)     |
| Coronary anatomy                            |                      | Male (v. female)                            | 1.29 (0.98–1.70)     |
| 1 vessel                                    | 1.00 (ref)           | Years since graduation (per additional yr) | 1.02 (1.00–1.05)     |
| 2 vessels                                   | 1.45 (1.34–1.57)     | Total consultations billed per year (per additional consultation) | 1.00 (1.00–1.10) |
| 3 vessels                                   | 1.66 (1.47–1.88)     | Total visits billed per yr (per additional visit) | 1.00 (1.00–1.10) |
| **Hospital**                                |                      | **Hospital**                                |                      |
| Annual catheterization volume (per additional 1000) | 1.00 (0.98–1.02) | **CABG, PCI and catheterization**            | 1.00 (ref)           |
| Hospital type                               |                      | Catheterization only                        | 1.05 (0.81–1.36)     |
| CABG, PCI and catheterization               | 1.00 (ref)           | PCI and catheterization only                | 0.86 (0.62–1.20)     |

Note: CABG = coronary artery bypass grafting, CCS = Canadian Cardiovascular Society, CI = confidence interval, COPD = chronic obstructive pulmonary disease, ECG = electrocardiogram, MI = myocardial infarction, OR = odds ratio, PCI = percutaneous coronary intervention, ref = reference group.

*Significant stenosis ≥ 70% (≥ 50% in left main coronary artery).

†Procedural physician.
tically significant difference in mortality between the medium and low revascularization ratio categories (HR 1.03, 95% CI 0.90–1.18). When we compared hospital categories within each treatment strategy, we found that patients who underwent PCI at a centre with a high revascularization ratio had a higher risk of death than those whose PCI was performed at a centre with a low revascularization ratio (HR 1.30, 95% CI 1.12–1.52). In contrast, mortality did not differ significantly between hospital categories among patients who underwent CABG or among those who received medical therapy alone (Table 4).

We found no significant differences in the risk of readmission with myocardial infarction between the 3 hospital categories (Table 4). When we compared treatment strategies, we found that patients who underwent PCI were less likely (HR 0.57, 95% CI 0.39–0.84), and patients who had CABG were more likely (HR 2.04, 95% CI 1.36–3.08), to be readmitted with myocardial infarction than patients who received medical therapy alone. (The full multivariable Cox models for all-cause mortality and myocardial infarction are available in Appendices 4 and 5, respectively, at www.cmaj.ca/lookup/suppl/doi:10.1503/cmaj.141372/-/DC1.)

**Interpretation**

In this study of initial treatment strategies for stable ischemic heart disease after coronary angiography, we found a twofold variation in revascularization ratios across the 18 cardiac centres in Ontario. Our analysis suggests that the variation was attributable primarily to differences in patient factors (67.4% of the variation). However, one-third of the variation remained unaccounted for. Importantly, the variation we observed between hospitals was associated with a small but statistically significant difference in all-cause mortality and myocardial infarction are available in Appendices 4 and 5, respectively, at www.cmaj.ca/lookup/suppl/doi:10.1503/cmaj.141372/-/DC1.)

A better understanding of both the drivers and consequences of variation is needed. We found that patient characteristics were the main driver of between-hospital variation in initial treatment strategy. The specific physician and hospital factors that we evaluated did not have a substantial impact on the variation, which is in contrast to findings from other studies on revascularization practice patterns and on the degree of optimal medical therapy. Potential explanations for our contrasting results are that patient factors are weighed more heavily in Ontario on the decision of initial treatment strategy for stable ischemic heart disease. Alternatively, given that 32.6% of the between-hospital variation was not accounted for in our study, practice patterns may have been

| Outcome                  | Hospital revascularization ratio | Adjusted HR* (95% CI) |
|--------------------------|---------------------------------|-----------------------|
| **All-cause mortality**  |                                 |                       |
| By revascularization ratio | High v. low                     | 1.12 (1.03–1.21)      |
|                          | Medium v. low                   | 1.03 (0.90–1.18)      |
| By treatment†            |                                 |                       |
| PCI v. medical therapy    | 0.84 (0.57–1.25)                |                       |
| CABG v. medical therapy   | 0.96 (0.62–1.47)                |                       |
| By treatment × revascularization ratio‡ |             |                       |
| CABG             | High v. low                     | 1.03 (0.86–1.23)      |
|                  | Medium v. low                   | 0.84 (0.67–1.05)      |
| PCI              | High v. low                     | 1.30 (1.12–1.52)      |
|                  | Medium v. low                   | 1.09 (0.94–1.27)      |
| Medical therapy     | High v. low                     | 1.06 (0.96–1.17)      |
|                    | Medium v. low                   | 1.10 (0.92–1.30)      |
| **Readmission with myocardial infarction** | | |
| By revascularization ratio | High v. low                     | 1.03 (0.84–1.26)      |
|                          | Medium v. low                   | 0.90 (0.72–1.13)      |
| By treatment†            |                                 |                       |
| PCI v. medical therapy    | 0.57 (0.39–0.84)                |                       |
| CABG v. medical therapy   | 2.04 (1.36–3.08)                |                       |
| By treatment × revascularization ratio‡ |             |                       |
| CABG                     | High v. low                     | 0.92 (0.72–1.19)      |
|                        | Medium v. low                   | 0.76 (0.58–1.00)      |
| PCI                      | High v. low                     | 1.18 (0.89–1.56)      |
|                        | Medium v. low                   | 0.93 (0.68–1.27)      |
| Medical therapy          | High v. low                     | 0.93 (0.74–1.18)      |
|                        | Medium v. low                   | 0.93 (0.73–1.19)      |

Note: CABG = coronary artery bypass grafting, CI = confidence interval, HR = hazard ratio, PCI = percutaneous coronary intervention.
*Adjusted for all patient, physician and hospital factors.
†Time-varying covariate for treatment received.
‡Analyzed by an interaction model.
influenced by components of hospital and physician culture that were not captured by the administrative data used in our analyses. Future research should explore other aspects of physician and hospital culture.

Because our study population had stable ischemic heart disease, overall mortality was low. Regardless, we observed small but statistically significant differences in mortality between hospitals with high and low revascularization ratios. It is important to note that our primary focus was not on the comparison of PCI or CABG with medical therapy. Although we did not find any differences in mortality, we found a statistically significant difference in readmission for myocardial infarction that requires further study. Rather, our primary focus was on the outcomes of all patients (having accounted for the actual treatment received) at high versus medium versus low revascularization ratio hospitals. We can only speculate as to the reason for the differences in outcomes we observed. One possible reason for the increased mortality at hospitals with a high revascularization ratio may be because these hospitals treat sicker patients; therefore, higher mortality was due to residual confounding from incomplete adjustment. We cannot discount this possibility, but we have 2 counterarguments. First, hospitals with a high revascularization ratio likely perform both higher risk PCI and CABG; as such, the lack of a significant between-hospital difference in mortality in the CABG group suggests against residual confounding as the primary driver. Second, the baseline characteristics of patients were relatively similar across the 3 hospital categories; therefore, the quality of care may have been lower at hospitals with a high revascularization ratio. This may seem contradictory to previous work that showed a strong association between procedural volume and quality of care.23-25

We did not compare centres according to their procedure volumes. Instead, we compared outcomes based on the threshold to revascularize a reference patient. Our results suggest that treatment at hospitals with a high revascularization ratio, which have a lower threshold for revascularization in a reference patient, is associated with increased mortality. We hypothesize that a lower threshold for revascularization in stable ischemic heart disease may put patients at risk of an invasive procedure without a substantial benefit. In addition, a focus on revascularization may divert attention away from aggressive medical therapy and thus prevent optimal medical therapy to be realized. We postulate that the reason our findings were isolated to PCI patients is because of the less discretionary nature of CABG,26 which would reduce the likelihood of inappropriate procedures.

Limitations
Several limitations to our study merit discussion. First, we limited our outcome measures to death and readmission with myocardial infarction. Because revascularization is primarily for symptom relief, the between-hospital variation we observed may have had an impact on quality of life or productivity, which we were unable to assess. Second, we were limited to evaluating physician and hospital factors that were captured in the databases available. Third, we conducted a complete case analysis, excluding 2% of patients in the cohort who had missing data. We do not anticipate that excluding these few patients from our analyses would qualitatively affect our conclusions. Finally, given the observational nature of the study, our findings should be considered hypothesis generating, not conclusive.

Conclusion
We identified important factors associated with the choice of initial treatment strategy for stable ischemic heart disease. Most of the variation in revascularization ratios across the 18 cardiac centres was explained by patient factors. Nonetheless, the variation was associated with potentially important differences in clinical outcomes.

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