Supplementary Online Content

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This supplementary material has been provided by the authors to give readers additional information about their work.

Section A: Research Design

- ☑ Does the study limit its scope to the assessment of disease conditions and/or procedures in an inpatient setting?
- ☑ Does the study infer that captured encounters do not represent distinct patients, but rather hospitalization records?
- ☑ Does the study avoid diagnosis/procedure-specific volume assessments for units that are not a part of the sampling frame of the NIS, and are therefore not representatively sampled, including
 - a) geographic units, like U.S. states
 - b) healthcare facilities (after 2011)
 - c) individual healthcare providers?

Section B: Data interpretation

- ☑ Does the study attempt to identify disease conditions or procedures of interest using administrative codes or their combinations that have been previously validated?
- ☑ Does the study limit its assessment to only in-hospital outcomes, rather than those occurring after discharge?
- ☑ Does the study limit assessment of outcomes to those accurately captured in the NIS, and avoid extrapolating non-specific secondary diagnoses to infer in-hospital events?

Section C: Data Analysis

- ☑ Does the study clearly account for the survey design of the NIS and its components clustering, stratification, and weighting?
- ☑ Does the study adequately address changes in data structure over time (for trend analyses)?

eTable 1: National Inpatient Sample (NIS) study checklist.

eTable 2: Elixhauser comorbidities used in the risk-adjustment model in this study.

| Heart failure | Peripheral vascular disease |
|-------------------------------------|--|
| Hypertension | Uncomplicated diabetes |
| Diabetes with chronic complications | Chronic pulmonary disease |
| Pulmonary circulation disease | Obesity |
| Weight loss | Paralysis |
| Other neurological disorders | Hypothyroidism |
| Renal failure | Fluid and electrolyte disorders |
| Liver disease | Alcohol abuse |
| Drug abuse | Psychoses |
| Depression | Peptic ulcer disease with bleeding |
| Chronic blood loss anemia | Iron deficiency anemia |
| Acquired immunodeficiency syndrome | Lymphoma |
| Metastatic cancer | Solid tumor without metastasis |
| Coagulopathy | Rheumatoid arthritis/collagen vascular disease |

eTable 3: Correlation between procedure and unadjusted in-hospital mortality

| Procedure | Spearman's rho | P-value |
|----------------|----------------|---------|
| | | |
| SAVR | -0.05 | 0.16 |
| SAVR + CABG | 0.09 | 0.01 |
| | 0.00 | . 0001 |
| MV repair | 0.33 | <.0001 |
| MV replacement | 0.13 | 0.0007 |

eTable 4: Trends in unadjusted median in-hospital mortality (and interquartile range) across deciles of procedure volume for isolated surgical aortic valve replacement (SAVR).

| SAVR | SAVR | Mortality for isolated SAVR, % (unadjusted) | | | | | |
|--------|---------------|---|-------------------|-------------------|---------|---------|--|
| Decile | Decile volume | | Lower Quartile | Upper Quartile | Minimum | Maximum | |
| 1 | 2-13 | 6.2 | 0.0 | 11.4 | 0.0 | 50.0 | |
| 2 | 13-20 | 4.2 | 0.0 | 7.7 | 0.0 | 16.7 | |
| 3 | 20-27 | 3.2 | 0.0 | 6.3 | 0.0 | 29.4 | |
| 4 | 27-33 | 3.8 | 1.9 | 7.1 | 0.0 | 15.7 | |
| 5 | 33-43 | 3.2 | 1.5 | 6.3 | 0.0 | 14.8 | |
| 6 | 43-53 | 2.9 | 2.2 | 5.3 | 0.0 | 17.1 | |
| 7 | 53-68 | 3.7 | 1.0 | 5.3 | 0.0 | 15.7 | |
| 8 | 68-87 | 3.8 | 1.7 | 5.6 | 0.0 | 11.4 | |
| 9 | 57-136 | 3.7 | 2.3 | 5.2 | 0.0 | 12.5 | |
| 10 | 136-1448 | 3.0 | 2.0 | 4.3 | 0.0 | 6.8 | |

eTable 5: Trends in unadjusted median in-hospital mortality (and interquartile range) across deciles of procedure volume for surgical aortic valve replacement (SAVR) + coronary artery bypass grafting (CABG).

| SAVR | SAVR | Mortality for SAVR + CABG, % (unadjusted) | | | | | |
|--------|---------------|---|-------------------|-------------------|---------|---------|--|
| Decile | Decile volume | | Lower Quartile | Upper Quartile | Minimum | Maximum | |
| 1 | 2-13 | 0.0 | 0.0 | 11.8 | 0.0 | 100.0 | |
| 2 | 13-20 | 0.0 | 0.0 | 10.0 | 0.0 | 50.0 | |
| 3 | 20-27 | 0.0 | 0.0 | 9.1 | 0.0 | 66.7 | |
| 4 | 27-33 | 4.0 | 0.0 | 9.1 | 0.0 | 26.1 | |
| 5 | 33-43 | 4.8 | 0.0 | 11.1 | 0.0 | 23.1 | |
| 6 | 43-53 | 4.5 | 0.0 | 8.3 | 0.0 | 25.0 | |
| 7 | 53-68 | 4.5 | 0.0 | 8.6 | 0.0 | 31.3 | |
| 8 | 68-87 | 3.6 | 0.0 | 7.7 | 0.0 | 16.7 | |
| 9 | 57-136 | 4.4 | 2.6 | 7.1 | 0.0 | 25.0 | |
| 10 | 136-1448 | 4.1 | 2.5 | 5.9 | 0.0 | 10.7 | |

eTable 6: Trends in unadjusted median in-hospital mortality (and interquartile range) across deciles of procedure volume for mitral valve (MV) repair.

| MV Repair | MV | Mortality for MV repair, % (unadjusted) | | | | | |
|------------------|------------------|---|-------------------|-------------------|---------|---------|--|
| Volume Decile | Repair volume | Median | Lower Quartile | Upper Quartile | Minimum | Maximum | |
| 1 | 1-2 | 0.0 | 0.0 | 0.0 | 0.0 | 50.0 | |
| 2 | 2-3 | 0.0 | 0.0 | 0.0 | 0.0 | 100.0 | |
| 3 | 3-5 | 0.0 | 0.0 | 0.0 | 0.0 | 66.7 | |
| 4 | 5-7 | 0.0 | 0.0 | 6.3 | 0.0 | 100.0 | |
| 5 | 7-9 | 0.0 | 0.0 | 7.1 | 0.0 | 33.3 | |
| 6 | 9-12 | 0.0 | 0.0 | 5.1 | 0.0 | 16.7 | |
| 7 | 12-16 | 0.0 | 0.0 | 8.3 | 0.0 | 28.6 | |
| 8 | 16-23 | 0.0 | 0.0 | 4.3 | 0.0 | 11.8 | |
| 9 | 24-39 | 2.3 | 0.0 | 4.8 | 0.0 | 16.0 | |
| 10 | 40-657 | 1.9 | 0.0 | 3.3 | 0.0 | 10.5 | |

| eTable 7: Trends | in unadjusted med | dian in-hospital | mortality | (and interquartile | range) |
|-------------------|-------------------|------------------|-----------|--------------------|--------|
| across deciles of | procedure volume | for mitral valve | (MV) rep | lacement. | |

| MV Replaceme | MV Replaceme | Mortality for MV replacement, % (unadjusted) | | | | | |
|---------------------|-----------------|--|-------------------|-------------------|---------|---------|--|
| nt Volume Decile | nt volume | Median | Lower Quartile | Upper Quartile | Minimum | Maximum | |
| 1 | 1-3 | 0.0 | 0.0 | 20.0 | 0.0 | 100.0 | |
| 2 | 3-5 | 0.0 | 0.0 | 20.0 | 0.0 | 50.0 | |
| 3 | 5-7 | 0.0 | 0.0 | 16.7 | 0.0 | 40.0 | |
| 4 | 7-9 | 5.2 | 0.0 | 14.3 | 0.0 | 40.0 | |
| 5 | 9-13 | 3.8 | 0.0 | 11.1 | 0.0 | 28.6 | |
| 6 | 13-16 | 6.3 | 0.0 | 10.0 | 0.0 | 27.3 | |
| 7 | 16-19 | 6.3 | 0.0 | 8.3 | 0.0 | 21.4 | |
| 8 | 19-25 | 6.6 | 3.4 | 10.3 | 0.0 | 23.1 | |
| 9 | 25-40 | 5.6 | 2.5 | 9.5 | 0.0 | 33.3 | |
| 10 | 40-398 | 5.5 | 2.9 | 8.0 | 0.0 | 25.8 | |

eTable 8: Risk-standardized mortality rates (median and interquartile range) for operative procedures in this study

| Procedures | Median RSMR | Interquartile range |
|-----------------|-------------|---------------------|
| Isolated SAVR | 2.9 | 2.8 to 3.2 |
| SAVR + CABG | 4.8 | 4.5 to 5.6 |
| MV repair* | 2.7 | 2.6 to 3.0 |
| MV replacement* | 6.4 | 6.1 to 7.2 |

* with and without CABG

eTable 9: Association between volume and in-hospital mortality in the hierarchical riskadjustment models for valve procedures, reported for a 10-unit increase in procedure volumes.

| | Beta-coefficient for 10- | Odds ratio for 10- | |
|-----------------|--------------------------|--------------------|---------|
| | unit increase in | unit increase in | P-value |
| | hospital procedure | hospital procedure | F-value |
| | volume | volume | |
| SAVR, isolated | -0.00080 | 0.992 | <.001 |
| SAVR + CABG | -0.00078 | 0.992 | <.001 |
| MV repair* | -0.00531 | 0.948 | <.001 |
| MV replacement* | -0.00763 | 0.927 | <.001 |

* with and without CABG

eTable 10: SAVR patient characteristics, 2011 to 2014

| | Calendar-years | | | | |
|--------------------------------------|----------------|------------|------------|------------|--|
| Characteristics | 2011 | 2012 | 2013 | 2014 | |
| Unweighted number | 12476 | 11309 | 11694 | 11619 | |
| Mainhted number (standard deviation) | 57950 | 56545 | 58470 | 58095 | |
| weighted number (standard deviation) | (5107) | (2031) | (2078) | (2044) | |
| Patient characteristics | | | | | |
| Female | 38.1 (0.6) | 36.4 (0.5) | 35.4 (0.4) | 34.4 (0.5) | |
| Race | | | | | |
| White | 75.8 (2.3) | 77.9 (1.3) | 78.9 (1.3) | 78.8 (1.2) | |
| Black | 4.4 (0.5) | 4.7 (0.3) | 4.5 (0.3) | 4.7 (0.3) | |
| Other | 10.1 (1.1) | 11.1 (1.0) | 9.9 (0.7) | 10.1 (0.7) | |
| Missing/Unknown | 9.7 (2.3) | 6.3 (1.0) | 6.8 (1.2) | 6.4 (1.0) | |
| Comorbid conditions | | | | | |
| Heart failure | 1.6 (0.2) | 1.4 (0.1) | 1.3 (0.1) | 1.4 (0.1) | |
| Valvular Heart Disease | 0.9 (0.1) | 0.8 (0.1) | 1.0 (0.1) | 0.9 (0.1) | |
| Peripheral Vascular Disease | 13.5 (0.9) | 13.0 (0.4) | 13.3 (0.5) | 12.7 (0.4) | |
| Chronic pulmonary disease | 18.4 (0.5) | 18.4 (0.4) | 18.4 (0.4) | 18.3 (0.4) | |
| Pulmonary circulation disease | 0.3 (0.05) | 0.5 (0.1) | 0.3 (0.1) | 0.3 (0.05) | |
| Hypertension | 67.3 (0.6) | 69.5 (0.5) | 68.7 (0.6) | 67.9 (0.5) | |
| Obesity | 15.3 (0.8) | 16.6 (0.5) | 18.2 (0.5) | 19.3 (0.5) | |
| Weight loss | 4.6 (0.4) | 4.6 (0.3) | 4.6 (0.3) | 3.9 (0.3) | |
| Paralysis | 1.5 (0.1) | 1.7 (0.1) | 1.5 (0.1) | 1.3 (0.1) | |
| Other neurological disorders | 3.9 (0.2) | 4.2 (0.2) | 4.3 (0.2) | 4.3 (0.2) | |
| Diabetes w/o complications | 23.8 (0.7) | 25.9 (0.5) | 25.3 (0.5) | 24.0 (0.5) | |
| Diabetes w/ chronic complications | 4.4 (0.6) | 4.1 (0.3) | 4.7 (0.3) | 5.2 (0.3) | |
| Hypothyroidism | 10.9 (0.3) | 10.9 (0.3) | 11.4 (0.3) | 11.0 (0.3) | |
| Renal failure | 15.3 (0.6) | 14.1 (0.4) | 13.2 (0.4) | 14.9 (0.3) | |
| Fluid and electrolyte disorders | 30.6 (1.7) | 31.8 (1.1) | 32.6 (1.0) | 34.1 (0.9) | |
| Liver disease | 1.4 (0.1) | 1.4 (0.1) | 1.4 (0.1) | 1.9 (0.1) | |
| Alcohol abuse | 1.8 (0.1) | 2.0 (0.1) | 2.4 (0.1) | 2.4 (0.1) | |
| Drug abuse | 0.9 (0.1) | 1.0 (0.1) | 1.2 (0.1) | 1.2 (0.1) | |
| Psychoses | 1.6 (0.2) | 1.8 (0.1) | 1.9 (0.1) | 1.9 (0.1) | |
| Depression | 4.9 (0.3) | 5.6 (0.2) | 5.2 (0.2) | 6.1 (0.2) | |
| Chronic blood loss anemia | 1.5 (0.3) | 1.0 (0.1) | 1.0 (0.1) | 0.9 (0.1) | |
| Peptic ulcer disease | 0.0 (0.01) | 0.0 (0.01) | 0.0 (0.01) | 0.0 (0.02) | |
| Iron deficiency anemia (ANEMDEF) | 16.9 (1.5) | 15.2 (0.6) | 14.6 (0.6) | 13.2 (0.5) | |
| AIDS | 0.1 (0.02) | 0.1 (0.03) | 0.1 (0.02) | 0.1 (0.02) | |
| Lymphoma | 0.7 (0.1) | 0.7 (0.1) | 0.7 (0.1) | 0.7 (0.1) | |
| Metastatic cancer | 0.2 (0.05) | 0.2 (0.04) | 0.1 (0.03) | 0.2 (0.04) | |
| Solid tumor without metastasis | 1.2 (0.1) | 1.2 (0.1) | 1.1 (0.1) | 1.0 (0.1) | |
| Coagulopathy | 27.5 (1.4) | 28.6 (0.9) | 30.3 (0.9) | 30.7 (0.8) | |
| Arthritis | 2.5 (0.2) | 2.7 (0.2) | 2.6 (0.1) | 2.5 (0.1) | |
| Death | 3.2 (0.2) | 3.0 (0.2) | 2.8 (0.2) | 2.9 (0.2) | |

Numbers represent % and SE unless otherwise specified

Abbreviations: SD – standard deviation, SEM – standard error of mean, C.I. – confidence interval, ESRD – End-stage renal disease

eTable 11: MV repair and replacement patient characteristics, 2011 to 2014

| | Calendar-years | | | | |
|-----------------------------------|----------------|------------|------------|------------|--|
| Characteristics | 2011 | 2012 | 2013 | 2014 | |
| Unweighted number | 5546 | 5060 | 5304 | 5434 | |
| Weighted number | 25817 | 25300 | 26520 | 27170 | |
| weighted humber | (2296) | (1209) | (1253) | (1251) | |
| Patient characteristics | | | | | |
| Female | 47.1 (0.9) | 45.4 (0.8) | 46.5 (0.7) | 46.4 (0.7) | |
| Race | | | | | |
| White | 70.4 (2.4) | 72.8 (1.7) | 72.5 (1.5) | 71.8 (1.4) | |
| Black | 8.3 (0.7) | 8.7 (0.6) | 8.9 (0.5) | 9.1 (0.6) | |
| Other | 11.6 (1.5) | 12.3 (1.0) | 12.0 (0.8) | 12.1 (0.8) | |
| Missing/Unknown | 9.7 (2.4) | 6.2 (1.6) | 6.7 (1.4) | 7.0 (1.3) | |
| Comorbid conditions | | | | | |
| Heart failure | 2.4 (0.2) | 2.2 (0.2) | 2.5 (0.2) | 2.8 (0.2) | |
| Valvular Heart Disease | 1.6 (0.2) | 1.7 (0.2) | 1.3 (0.2) | 1.7 (0.2) | |
| Peripheral Vascular Disease | 7.3 (0.6) | 7.0 (0.4) | 6.5 (0.4) | 7.4 (0.4) | |
| Chronic pulmonary disease | 17.4 (0.8) | 17.4 (0.6) | 17.6 (0.6) | 17.9 (0.6) | |
| Pulmonary circulation disease | 0.7 (0.1) | 0.7 (0.1) | 0.7 (0.1) | 0.9 (0.1) | |
| Hypertension | 52.8 (0.8) | 55.1 (0.9) | 55.9 (0.8) | 54.9 (0.7) | |
| Obesity | 9.0 (0.6) | 10.2 (0.5) | 10.9 (0.6) | 12.2 (0.5) | |
| Weight loss | 6.6 (0.7) | 5.4 (0.4) | 6.3 (0.5) | 6.3 (0.5) | |
| Paralysis | 1.8 (0.2) | 2.1 (0.2) | 1.7 (0.2) | 1.8 (0.2) | |
| Other neurological disorders | 3.8 (0.3) | 4.2 (0.3) | 3.6 (0.2) | 4.2 (0.3) | |
| Diabetes w/o complications | 14.2 (0.8) | 14.3 (0.6) | 14.3 (0.6) | 13.9 (0.6) | |
| Diabetes w/ chronic complications | 2.6 (0.4) | 2.1 (0.2) | 3.4 (0.3) | 3.2 (0.3) | |
| Hypothyroidism | 9.6 (0.4) | 9.7 (0.4) | 10.4 (0.4) | 9.2 (0.4) | |
| Renal failure | 13.1 (0.6) | 12.1 (0.6) | 11.8 (0.5) | 13.2 (0.5) | |
| Fluid and electrolyte disorders | 32.8 (1.8) | 33.4 (1.5) | 34.9 (1.2) | 38.6 (1.3) | |
| Liver disease | 1.6 (0.2) | 1.1 (0.1) | 1.7 (0.2) | 1.3 (0.1) | |
| Alcohol abuse | 2.0 (0.2) | 2.1 (0.2) | 2.0 (0.2) | 2.3 (0.2) | |
| Drug abuse | 1.3 (0.2) | 1.6 (0.2) | 1.6 (0.2) | 2.0 (0.2) | |
| Psychoses | 2.1 (0.3) | 1.8 (0.2) | 2.3 (0.2) | 2.2 (0.2) | |
| Depression | 6.4 (0.4) | 5.6 (0.3) | 6.2 (0.3) | 5.9 (0.3) | |
| Chronic blood loss anemia | 1.7 (0.3) | 0.7 (0.1) | 0.8 (0.1) | 0.9 (0.2) | |
| Peptic ulcer disease | 0 (0) | 0.0 (0.02) | 0 (0) | 0.0 (0.02) | |
| Iron deficiency anemia | 17.1 (1.9) | 14.1 (0.7) | 13.8 (0.7) | 12.4 (0.6) | |
| AIDS | 0.1 (0.04) | 0.1 (0.06) | 0.1 (0.05) | 0.1 (0.05) | |
| Lymphoma | 0.5 (0.1) | 0.6 (0.1) | 0.6 (0.1) | 0.4 (0.1) | |
| Metastatic cancer | 0.1 (0.04) | 0.2 (0.06) | 0.1 (0.05) | 0.2 (0.1) | |
| Solid tumor without metastasis | 0.8 (0.1) | 0.8 (0.1) | 0.6 (0.1) | 0.6 (0.1) | |
| Coagulopathy | 25.7 (1.6) | 25.7 (1.0) | 25.6 (0.9) | 26.6 (0.8) | |
| Arthritis | 2.2 (0.2) | 2.5 (0.2) | 2.6 (0.2) | 2.6 (0.2) | |
| Death | 3.8 (0.3) | 3.9 (0.3) | 3.7 (0.3) | 3.6 (0.3) | |

Numbers represent % and SE unless otherwise specified

Abbreviations: SD – standard deviation, SEM – standard error of mean, C.I. – confidence interval, ESRD – End-stage renal disease

eFigure 1: Patient selection flowsheet



Abbreviations: SAVR: Surgical Aortic Valve Replacement, MV: Mitral valve, CABG: Coronary Artery Bypass

Grafting

eFigure 2: Unadjusted in-hospital mortality by hospital volume. Horizontal line represents the mean mortality for each condition, the dotted lines represent the 95% confidence intervals as a binominal distribution around the mean volume for the procedure. The values between the confidence intervals represent those expected by chance alone.









