

Supplementary Information

Article title

Frailty Index based on laboratory tests and in-hospital falls among older adults

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Supplementary Table 1

Original version of STRATIFY and the modified versions used in the present study.

| | Item | Original STRATIFY* | Modified STRATIFY |
|---|-----------------------|--|--|
| 1 | History of falls | Did the patient present to hospital with a fall or has he or she fallen on the ward since admission? (Yes = 1, No = 0) | History of falls within 1 year.‡ (Yes = 1, No = 0) |
| 2 | Mental status | Agitated?† (Yes = 1, No = 0) | Is the patient confused, disoriented, or agitated?†.§ (Yes = 1, No = 0) |
| 3 | Vision | Visually impairment to the extent that everyday function is affected?† (Yes = 1, No = 0) | (Same as the original) |
| 4 | Toileting | In need of especially frequent toileting?† (Yes = 1, No = 0) | (Same as the original) |
| 5 | Transfer and mobility | Transfer and mobility score of 3 or 4?† (Yes = 1, No = 0) | (Same as the original) |

Notes: STRATIFY = St. Thomas's Risk Assessment Tool in Falling Elderly Inpatients

*Based on Oliver et al. BMJ 1997; 315: 1049.

†Judged by nurses. No attempt was made to standardize the formation of these judgments.

‡Based on history-taking.

§Referring to Papaioannou et al. BMC Medicine 2004; 2: 1.

Supplementary Table 2

Laboratory data used to construct the Frailty Index-laboratory (FI-lab).

| Variable* | Unit | Total population | | Male | | Female | |
|-------------------|----------------------|------------------|-----------------|---------------|-----------------|---------------|-----------------|
| | | 0 (Normal) | 1 (Abnormal) | 0 (Normal) | 1 (Abnormal) | 0 (Normal) | 1 (Abnormal) |
| WBC | ×10 ³ /μL | 3.3-8.6 | <3.3 or >8.6 | | | | |
| Neutrophil | ×10 ³ /μL | 1.0-7.5 | <1.0 or >7.5 | | | | |
| Lymphocyte | ×10 ³ /μL | 1.0-4.0 | <1.0 or >4.0 | | | | |
| Hemoglobin | g/dL | | | 13.7-16.8 | <13.7 or >16.8 | 11.6-14.8 | <11.6 or >14.8 |
| Hematocrit | % | | | 40.7-50.1 | <40.7 or >50.1 | 35.1-44.4 | <35.1 or >44.4 |
| MCV | fL | 83.6-98.2 | 83.6< or >98.2 | | | | |
| RDW | % | 11-14 | <11 or >14 | | | | |
| Platelet count | ×10 ³ /μL | 158-348 | <158 or >348 | | | | |
| Total protein | g/dL | 6.6-8.1 | <6.6 or >8.1 | | | | |
| Albumin | g/dL | 4.1-5.1 | <4.1 or >5.1 | | | | |
| Glucose | mg/dL | 73-109 | <73 or >109 | | | | |
| Urea | mg/dL | 8-20 | <8 or >20 | | | | |
| Creatinine | mg/dL | | | 0.65-1.07 | <0.65 or >1.07 | 0.46-0.79 | <0.46 or >0.79 |
| Uric acid | mg/dL | | | 3.7-7.8 | <3.7 or >7.8 | 2.6-5.5 | <2.6 or >5.5 |
| Sodium | mmol/L | 138-145 | <138 or >145 | | | | |
| Potassium | mmol/L | 3.6-4.8 | <3.6 or >4.8 | | | | |
| Chloride | mmol/L | 101-108 | <101 or >108 | | | | |
| Corrected calcium | mg/dL | 8.8-10.1 | <8.8 or >10.1 | | | | |
| AST | U/L | 13-30 | <13 or >30 | | | | |
| ALT | U/L | | | 10-42 | <10 or >42 | 7-23 | <7 or >23 |
| LDH | U/L | 124-222 | <124 or >222 | | | | |
| Total bilirubin | mg/dL | 0.4-1.5 | <0.4 or >1.5 | | | | |
| ALP (JSCC) | U/L | 106-322 | <106 or >322 | | | | |
| γGTP | U/L | | | 13-64 | <13 or >64 | 9-32 | <9 or >32 |
| Amylase | U/L | 44-132 | <44 or >132 | | | | |
| CK | U/L | | | 59-248 | <59 or >248 | 41-153 | <41 or >153 |
| Total cholesterol | mg/dL | 142-248 | <142 or >248 | | | | |
| Triglyceride | mg/dL | | | 40-234 | <40 or >234 | 30-117 | <30 or >117 |
| HDL-C | mg/dL | | | 38-90 | <38 or >90 | 48-103 | <48 or >103 |
| LDL-C | mg/dL | 65-163 | <65 or >163 | | | | |
| HbA1c | % | 4.9-6.0 | <4.9 or >6.0 | | | | |
| CRP | mg/dL | ≤0.14 | >0.14 | | | | |
| PT | % | 80-120 | <80 or >120 | | | | |
| APTT | % | 80-120 | <80 or >120 | | | | |
| Fibrinogen | mg/dL | 200-400 | <200 or >400 | | | | |

Notes: ALT = alanine transaminase; ALP = alkaline phosphatase; APTT = activated partial thromboplastin time; AST = aspartate transaminase; CK = creatine kinase; CRP = C-reactive protein; γGTP = gamma-glutamyl transpeptidase; HbA1c = hemoglobin A1c; HDL-C = high-density lipoprotein cholesterol; JSCC = Japan Society of Clinical Chemistry; LDH = lactate dehydrogenase; LDL-C = low-density lipoprotein-cholesterol; MCV = mean cell volume; PT = prothrombin time; RDW = red cell distribution width; WBC = white blood cells

*Cut-off values were obtained from our Department of Laboratory Medicine.

Supplementary Table 3

Items included in the standard non-laboratory Frailty Index (FI-clinical).

| Variable | Deficit value | | | | |
|---------------------------------|--------------------------------|-----------------|------------------------|------------------|-------------------------------------|
| | 0 points | 0.33 points | 0.5 points | 0.66 points | 1 point |
| 1 History of falls | absent | | | | present |
| 2 Muscle weakness | absent | | | | present |
| 3 Difficulty with balance | absent | | | | present |
| 5 Shuffling or festinating gait | absent | | | | present |
| 4 Contracture | absent | | | | present |
| 6 Frequent urination | absent | | | | present |
| 7 Nocturia | absent | | | | present |
| 8 BMI | 21.5-25.0 kg/m ² | | | | <21.5 or >25.0 kg/m ² |
| 9 Coronary artery disease | absent | | | | present |
| 10 Congestive heart failure | absent | | | | present |
| 11 Peripheral artery disease | absent | | | | present |
| 12 Dementia | absent | | | | present |
| 13 COPD | absent | | | | present |
| 14 Peptic ulcer disease | absent | | | | present |
| 15 Chronic kidney disease | absent | | | | present |
| 16 Cerebral vascular accident | absent | | | | present |
| 17 Diabetes mellitus | absent | | | | present |
| 18 Hypertension | absent | | | | present |
| 19 Use of sleep medication(s) | no | | | | yes |
| 20 Feeding | independent | | needs help | | totally dependent |
| 21 Position change in bed | independent | | needs help | | totally dependent |
| 22 Maintaining sitting position | independent | | | | totally dependent |
| 23 Sitting adjustment | independent | | | | totally dependent |
| 24 Transfer | independent | minor help | | major help | totally dependent |
| 25 Walking | independent | walks with help | | wheelchair bound | Immobile |
| 26 Use of walking aids | no | | | | yes |
| 27 Stairs | independent | | needs help | | total dependent |
| 28 Grooming | independent | | | | needs help with personal care |
| 29 Bathing | independent | | | | total dependent |
| 30 Dressing | independent | | needs help | | total dependent |
| 31 Toilet | independent | | needs help | | totally dependent |
| 32 Bowel control | continent | | occasional accident | | incontinent |
| 33 Use of diaper for defecation | no | | | | yes |
| 34 Bladder control | continent | | occasional accident | | incontinent |
| 35 Use of diaper for urination | no | | | | yes |
| 36 Disorientation | absent | | | | present |
| 37 Impaired comprehension | absent | | | | present |
| 38 Impaired memory | absent | | | | present |
| 39 Swallowing difficulty | absent | | | | present |
| 40 Hearing problems | no problems | | | | hear only loud voices |

Supplementary Table 4

Fall risk assessment tool used at the authors' hospital.

| Major category | Items |
|---|---|
| 1. History of falls | <ol style="list-style-type: none">1. History of falls2. History of syncope3. History of convulsions4. History of epilepsy |
| 2. Disorder of the sensory system | <ol style="list-style-type: none">1. Visual impairment2. Hearing impairment |
| 3. Disorder of the musculoskeletal system | <ol style="list-style-type: none">1. Muscle weakness2. Paralysis, numbness3. Dizziness4. Lightheadedness5. Joint abnormalities/contractures/deformations6. Gait disturbance |
| 4. Disorder of cognition function | <ol style="list-style-type: none">1. Disturbance of consciousness2. Restlessness3. Disorientation4. Memory disturbance5. Impaired judgment6. Not able to press a buzzer |
| 5. Dysfunction of urination/defecation | <ol style="list-style-type: none">1. Needs helper2. Fecal incontinence3. Frequent urination4. Go to bathroom often at night5. Needs diapers for defecation6. Urinary incontinence7. Needs diapers for urination |
| 6. Mobility dysfunction/dependency | <ol style="list-style-type: none">1. Inability to stand or walk2. Cane3. Walker4. Wheelchair5. Need assistance6. Use of intravenous infusion, placement of drains, oxygenation |
| 7. Medication | <ol style="list-style-type: none">1. Hypnotics (occasionally)2. Hypnotics (every day)3. Psychotropic drugs4. Anticancer agents5. Opioids6. Analgesics7. Antihypertensive agents |

Supplementary Table 5

Common primary reasons for admission.

| Rank | Major diagnostic category | n | %* |
|------|---|------|------|
| 1 | Cancer and complications | 2650 | 44.3 |
| 2 | Arrhythmia | 263 | 4.4 |
| 3 | Heart failure | 212 | 3.5 |
| 4 | Arterial diseases | 189 | 3.2 |
| 5 | Neurological diseases other than stroke | 175 | 2.9 |
| 6 | Benign gastrointestinal tumor | 156 | 2.6 |
| 7 | Renal failure | 130 | 2.2 |
| 8 | Biliary diseases | 130 | 2.9 |
| 9 | Cardiac valvular diseases | 110 | 1.8 |
| 10 | Diabetes mellitus and complications | 99 | 1.7 |
| 11 | Acute coronary syndrome | 82 | 1.4 |
| 12 | Pneumonia | 65 | 1.1 |
| 13 | Gastrointestinal ileus | 60 | 1.0 |
| 14 | Gastrointestinal bleeding | 57 | 1.0 |
| 15 | Autoimmune disorders | 45 | 0.8 |

*Percentage of cases to total admissions (n=5984).

Supplementary Table 6

Baseline characteristics of the study population and outcomes according to FI-lab score (n=5984).

| | Total | Low (<0.25) | Moderate (0.25–0.4) | High (>0.4) | p ^g |
|--|---------------|---------------|---------------------|--------------|--------------------|
| Participants, n (%) | 5984 (100.0) | 2383 (39.8) | 1859 (31.1) | 1742 (29.1) | |
| Age, years | 73.0 ± 7.3 | 71.5 ± 6.7 | 73.7 ± 7.3 | 74.4 ± 7.7 | <.001 ^h |
| Male sex | 3798 (63.5) | 1383 (58.0) | 1253 (67.4) | 1162 (66.7) | <.001 ⁱ |
| Emergency hospitalization | 1551 (25.9) | 236 (9.9) | 395 (21.2) | 920 (52.8) | <.001 ⁱ |
| Body mass index, kg/m ² | 22.3 ± 3.8 | 22.8 ± 3.5 | 22.3 ± 3.9 | 21.8 ± 4.0 | <.001 ^h |
| Barthel Index ^a , median (IQR) | 100 (100–100) | 100 (100–100) | 100 (100–100) | 100 (80–100) | <.001 ^h |
| Barthel Index ^a , mean ± SD ^b | 94.1 ± 15.5 | 98.0 ± 8.2 | 95.4 ± 13.3 | 87.4 ± 21.7 | |
| CCI ^c | 2 (1–2) | 2 (0–2) | 2 (1–2) | 2 (1–3) | <.001 ^h |
| Polypharmacy score ^d | 1 (0–2) | 1 (0–2) | 1 (0–2) | 1 (1–2) | <.001 ^h |
| STRATIFY score ^e , median (IQR) | 0 (0–1) | 0 (0–0) | 0 (0–1) | 0 (0–1) | <.001 ^h |
| STRATIFY score ^e , mean ± SD ^b | 0.42 ± 0.72 | 0.30 ± 0.56 | 0.39 ± 0.70 | 0.62 ± 0.89 | |
| History of falls | 807 (13.5) | 201 (8.4) | 255 (13.7) | 351 (20.1) | <.001 ⁱ |
| Mental state abnormality | 353 (5.9) | 76 (3.2) | 94 (5.1) | 183 (10.5) | <.001 ⁱ |
| Vision impairment | 442 (7.4) | 188 (7.9) | 117 (6.3) | 137 (7.9) | .83 ⁱ |
| Frequent urination | 419 (7.0) | 161 (6.8) | 133 (7.2) | 125 (7.2) | .59 ⁱ |
| Transferring and/or mobility impairment | 504 (8.4) | 80 (3.4) | 133 (7.2) | 291 (16.7) | <.001 ⁱ |
| FI-lab ^f | 0.31 ± 0.16 | 0.15 ± 0.06 | 0.32 ± 0.04 | 0.51 ± 0.09 | <.001 ^h |
| Measured ratio | 0.87 ± 0.08 | 0.88 ± 0.07 | 0.87 ± 0.08 | 0.87 ± 0.08 | .027 ^h |
| FI-clinical ^f | 0.11 ± 0.09 | 0.08 ± 0.06 | 0.11 ± 0.08 | 0.16 ± 0.12 | <.001 ^h |
| Outcomes | | | | | |
| In-hospital falls | 175 (2.9) | 42 (1.8) | 38 (2.0) | 95 (5.5) | <.001 ⁱ |
| In-hospital injurious falls | 32 (0.5) | 10 (0.4) | 7 (0.4) | 15 (0.9) | .07 ⁱ |
| In-hospital mortality | 95 (1.6) | 3 (0.1) | 8 (0.4) | 84 (4.8) | <.001 ⁱ |
| Length of hospital stay | 9 (5–16) | 8 (4–12) | 8 (4–15) | 12 (7–23) | <.001 ^h |
| Discharge home | 5382 (89.9) | 2271 (95.5) | 1718 (92.5) | 1393 (80.3) | <.001 ⁱ |

Notes: Data are presented as the mean ± standard deviation, median (interquartile range), or number (percentage). CCI = Charlson Comorbidity Index; FI-clinical = standard non-laboratory Frailty Index; FI-lab = Frailty Index based on laboratory tests; IQR = interquartile range; SD = standard deviation; STRATIFY = St. Thomas's Risk Assessment Tool in Falling Elderly Inpatients.

^aBarthel Index score ranges from 0 to 100, with a higher score indicating higher function.

^bThe mean and SD are shown for reference.

^cThe CCI score ranges from 0 to 37, with a higher score indicating more comorbidities.

^dThe polypharmacy score ranges from 0 to 6, with a higher score indicating more medications in use.

^eThe STRATIFY score ranges from 0 to 5, with a higher score indicating a higher fall risk.

^fThe FI-lab and FI-clinical scores range from 0 to 1, with a higher score indicating worse frailty.

^gTest for trends among the three subgroups.

^hJonckheere-Terpstra test.

ⁱCochran-Armitage test.

Supplementary Table 7

Results of subgroup analyses.

| Subgroup | n | In-hospital fall, n (%) | | ROC-AUC | | p ^b |
|----------|------|----------------------------|-------|----------|---|----------------|
| | | | | STRATIFY | STRATIFY including FI-lab ^s | |
| Overall | 5984 | 175 | (2.9) | 0.674 | 0.718 | <0.001 |
| Age | | | | | | |
| < 75 | 3461 | 82 | (2.4) | 0.672 | 0.721 | 0.014 |
| ≥ 75 | 2523 | 93 | (3.7) | 0.647 | 0.698 | 0.002 |
| Sex | | | | | | |
| Male | 3798 | 103 | (2.7) | 0.657 | 0.705 | 0.006 |
| Female | 2186 | 72 | (3.3) | 0.704 | 0.741 | 0.025 |

Notes: FI-lab = Frailty Index-laboratory; ROC-AUC = area under the receiver-operating characteristic curve; STRATIFY = St.

Thomas's Risk Assessment Tool in Falling Elderly Inpatients.

^aA total of 6 items: 5 STRATIFY items and 1 item on whether FI-lab is higher than 0.4.

^bComparison between STRATIFY and STRATIFY including FI-lab.

Supplementary Table 8

Results of sensitivity analyses.

| Model ^a | ROC-AUC | 95% CI | p |
|---|---------|---------------|--------|
| With or without polypharmacy score ^b | | | |
| STRATIFY | 0.674 | (0.632–0.716) | ref |
| STRATIFY + polypharmacy score ^b | 0.675 | (0.633–0.716) | 0.64 |
| STRATIFY + FI-lab | 0.715 | (0.677–0.753) | 0.018 |
| STRATIFY + FI-lab + polypharmacy score ^b | 0.715 | (0.677–0.753) | 0.017 |
| STRATIFY + FI-clinical | 0.707 | (0.667–0.747) | 0.003 |
| STRATIFY + FI-clinical + polypharmacy score ^b | 0.709 | (0.668–0.749) | 0.001 |
| Changing cut-off point of FI-lab | | | |
| FI-lab (binary variable: cut-off 0.4) | 0.659 | (0.616–0.702) | ref |
| FI-lab (binary variable: cut-off 0.345) | 0.670 | (0.628–0.712) | 0.34 |
| STRATIFY including FI-lab ^c (binary variable: cut-off 0.4) | 0.718 | (0.681–0.755) | ref |
| STRATIFY including FI-lab ^c (binary variable: cut-off 0.345) | 0.725 | (0.690–0.759) | 0.40 |
| Changing fall prediction tool | | | |
| Fall risk assessment tool in the author's hospital | 0.667 | (0.626–0.709) | ref |
| STRATIFY | 0.674 | (0.632–0.716) | 0.69 |
| Fall risk assessment tool including FI-lab ^d | 0.691 | (0.652–0.731) | <0.001 |

Notes: CI, confidence interval; FI-lab, Frailty Index-laboratory; ROC-AUC, area under the receiver-operating characteristic curve;

STRATIFY, St. Thomas's Risk Assessment Tool in Falling Elderly Inpatients.

^aAll models include age and sex as covariates.

^bThe polypharmacy score ranges from 0 to 6, with a higher score indicating more medications in use.

^cA total of 6 items: 5 STRATIFY items and 1 item on whether FI-lab is higher than the cut-off.

^dA total of 8 items: 7 items of fall risk assessment tool and 1 item on whether the FI-lab is higher than 0.4.

Supplementary Figure 1

Frequency distributions for the STRATIFY (A), standard non-laboratory Frailty Index (FI-clinical) (B), and Frailty Index-laboratory (FI-lab) (C) (n=5984). STRATIFY = St. Thomas's Risk Assessment Tool in Falling Elderly Inpatients

