

SEVERITY, NOT PRESENCE, OF DIASTOLIC DYSFUNCTION DRIVES PERIOPERATIVE CARDIOVASCULAR RISK BEFORE NON-CARDIAC SURGERY: A PROSPECTIVE COHORT STUDY PROPOSING A SEVERITY-FIRST TRIAGE FRAMEWORK.

Abstract:

Background: Diastolic dysfunction (DD) with preserved ejection fraction is prevalent before non-cardiac surgery, yet its prognostic weight remains contested. Guidelines flag DD without an actionable threshold. We hypothesized that DD severity, indexed by elevated left ventricular filling pressures (LVFP), drives perioperative risk.

Methods: This prospective single-center cohort enrolled 102 adults scheduled for elective non-cardiac surgery (March 2025 to February 2026). All underwent preoperative transthoracic echocardiography per 2016 ASE/EACVI criteria. Isolated DD was preserved left ventricular ejection fraction (LVEF) \geq 50% with abnormal diastolic function. The primary endpoint was in-hospital major adverse cardiovascular events (MACE), comparing isolated DD (n = 44) and normal controls (n = 36). Performance used logistic regression with bootstrap-corrected area under the curve (AUC).

Results: Of 80 LVEF-preserved patients, 55.0% had isolated DD, with heavier comorbidity (diabetes 59.1% vs. 16.7%, $P < 0.001$) and 7-fold higher median NT-proBNP (733 vs. 98 pg/ml, $P < 0.001$). MACE rates were similar (22.7% vs. 16.7%; odds ratio [OR] 1.47, 95% confidence interval [CI] 0.48 to 4.53). Binary DD added negligible discrimination over Lee plus NT-proBNP (Δ AUC +0.002); severe DD trended stronger (OR 3.22, 95% CI 0.79 to 13.19), lifting AUC from 0.70 to 0.77. Isolated DD prompted a 5-fold rise in preoperative optimization (50.0% vs. 11.1%, $P < 0.001$).

Conclusions: DD should be treated not as a binary alarm but as a graded signal whose severity, indexed by LVFP, drives prognosis. We propose a Severity-First Diastolic Triage framework integrating Lee score, NT-proBNP, and LVFP to rationalize preoperative echocardiography.

Key words:-

Diastolic heart failure; Echocardiography; Heart failure with preserved ejection fraction; Natriuretic peptide, brain; Noncardiac surgery; Perioperative care; Preoperative care; Risk assessment; Ventricular dysfunction, left.

Introduction:-

Heart failure with preserved ejection fraction (HFpEF) now accounts for approximately half of all incident heart failure cases worldwide, and its prevalence is rising with population aging and the global epidemic of cardiometabolic disease [1,2]. Left ventricular diastolic dysfunction (DD)—the morphological substrate of HFpEF—follows a pathophysiological continuum from isolated impaired relaxation to elevated filling pressures and overt clinical decompensation [3,4]. In community cohorts, DD affects 30% to 50% of adults over 60 years of age [5,6] and rises to 55% or more among patients presenting for elective non-cardiac surgery [7].

In the perioperative arena, this matters acutely. General anesthesia, surgical stress, and large fluid shifts impose hemodynamic loads that a stiff, non-compliant ventricle tolerates poorly: subendocardial ischemia, atrial fibrillation, pulmonary congestion, and demand-supply ischemic injury are all overrepresented in patients with abnormal diastolic function [8–10]. The recently published 2024 AHA/ACC/ACS/ASNC/HRS/SCA/SCCT/SCMR/SVM perioperative cardiovascular guideline [11] and the 2022 ESC/ESAIC guideline [12] both flag DD as a relevant prognostic marker, but neither specifies a clinical threshold for action, and routine preoperative transthoracic echocardiography (TTE) is explicitly not recommended [11,12].

Three problems have so far kept the field paralyzed. First, DD is rarely truly isolated in the published evidence base: most retrospective series mix patients with concomitant systolic dysfunction or hypertrophy, blurring its independent contribution [13,14]. Second, the very definition of DD has shifted across the 2009 and 2016 ASE/EACVI iterations, with a decisive move from the E/A ratio toward integrated assessment of left ventricular filling pressures (LVFP) [15]. Third, and most importantly, the prognostic gain of DD must now be tested incrementally against well-validated tools—the Revised Cardiac Risk Index (Lee score), NT-proBNP [16,17], and the Duke Activity Status Index [18]—rather than in isolation. The SOLOMON study (2023) demonstrated that even global longitudinal strain offers only modest incremental discrimination over established scores [19], reinforcing the need for parsimony when adding echocardiographic parameters to perioperative risk models.

A consequence of this paralysis is clinical: the simple mention of “diastolic dysfunction” on a preoperative TTE report continues to trigger disproportionate care escalations—cardiology consultation, surgery deferral, intensive care unit (ICU) pre-admission—in patients in whom the true incremental risk is small. Conversely,

55 severe DD with elevated LVFP, which does carry meaningful risk, is sometimes lost in the noise. The field
56 needs a disciplined, severity-graded approach that the anesthesiologist can apply at the bedside.

57 Against this background, we conducted a prospective single-center cohort study with three prespecified
58 objectives: (1) to characterize the prevalence and clinical phenotype of isolated DD with preserved LVEF in
59 consecutive non-cardiac surgical patients; (2) to quantify its association with cardiovascular comorbidities, NT-
60 proBNP elevation, and preoperative therapeutic decision-making; and (3) to test, against established predictors
61 (Lee score, NT-proBNP, surgical risk), the hypothesis that DD severity—captured by elevated LVFP—rather
62 than its binary presence, drives perioperative MACE risk. Building on our findings, we propose a Severity-First
63 Diastolic Triage (SFDT) framework intended to rationalize the preoperative echocardiographic decision
64 pathway, in line with the 2024 AHA/ACC [11] and 2023 ESC focused update on heart failure [20].

65 **Materials and Methods:-**

66 **Study design and setting**

67 This was a prospective single-center observational cohort study conducted at Oued Eddahab Military hospital
68 between 17 March 2025 and 1 February 2026. The protocol was approved by “Comité d’Ethique pour la
69 Recherche Biomédicale Université Mohammed V CERB – Rabat” with approval number 31/25 by the local
70 Institutional Review Board and complied with the Declaration of Helsinki (revised 2013) and the STROBE
71 statement for observational studies [21]. Written informed consent was obtained from every participant. The
72 study was not interventional; all clinical decisions remained the responsibility of the attending teams.

73 **Participants**

74 Consecutive adults (≥ 18 years of age) admitted for elective non-cardiac surgery and seen at the standardized
75 preoperative anesthesia consultation were eligible. Exclusion criteria were: emergency surgery; cardiac or
76 thoracic surgery; chronic renal replacement therapy; and inability to obtain interpretable echocardiographic
77 windows. No patient was excluded on the basis of age, comorbidity burden, or surgical risk class. The sex of all
78 participants was recorded. The study cohort included both male and female adults; no a priori sex-based
79 exclusion was applied.

80 **Echocardiographic assessment**

81 All TTE examinations were performed by a single experienced cardiologist blinded to clinical outcomes, using a
82 standard echocardiography system equipped with a 2.5–3.5 MHz transducer. Image acquisition and analysis
83 followed the 2016 ASE/EACVI recommendations [15].

84 LVEF was calculated by the modified biplane Simpson method; preserved LVEF was defined as $\geq 50\%$.
85 Diastolic function was classified per the 2016 ASE/EACVI algorithm integrating septal and lateral mitral
86 annular tissue Doppler velocities (e'), average E/e' ratio, indexed left atrial volume, and peak tricuspid
87 regurgitation velocity. Categories were: (1) normal; (2) isolated impaired relaxation (mild DD); (3) abnormal
88 compliance; and (4) combined relaxation and compliance abnormalities (advanced DD). LVFP was classified as
89 normal or elevated by the same algorithm; elevated LVFP defined severe DD.

90 Four echocardiographic phenotypes were prespecified by combining LVEF and diastolic function: (1) Normal—
91 preserved LVEF + normal diastolic function; (2) Isolated DD—preserved LVEF + abnormal diastolic function;
92 (3) Combined SD + DD—reduced LVEF + abnormal diastolic function; and (4) Isolated SD—reduced LVEF +
93 normal diastolic function. The primary analytic comparison contrasted Isolated DD with Normal (preserved-
94 LVEF cohort, $n = 80$).

95 **Clinical and laboratory variables**

96 Demographics, cardiovascular risk factors (diabetes, hypertension, dyslipidemia, obesity, smoking),
97 cardiovascular history, current medications, New York Heart Association (NYHA) dyspnea class, full six-item
98 Lee/Revised Cardiac Risk Index (RCRI) [22], surgical risk stratum per 2022 ESC criteria [12], and preoperative
99 NT-proBNP (within 30 days of surgery, by a single standardized electrochemiluminescence assay) were
100 recorded prospectively on a structured case-report form. The 300 pg/ml NT-proBNP threshold from the 2022
101 ESC/ESAIC guideline [12] was used as the principal cut-off, with sensitivity analyses at the 125 pg/ml
102 threshold endorsed by recent VISION substudies [16].

103 **Outcomes**

104 The primary outcome was a composite of in-hospital major adverse cardiovascular events (MACE), prespecified
105 as: all-cause death; non-fatal myocardial infarction (per the Fourth Universal Definition [23]); cardiogenic

106 shock; acute respiratory failure requiring ventilatory support; symptomatic arrhythmia requiring intervention; or
107 unplanned admission to the intensive or intermediate care unit. Secondary outcomes were: preoperative
108 therapeutic adjustments (initiation/modification of beta-blockers, diuretics, statins, antithrombotic agents, or
109 blood-pressure optimization); preoperative admission for medical optimization; intraoperative red-cell
110 transfusion; and length of hospital stay. Outcomes were adjudicated by two reviewers blinded to
111 echocardiographic phenotype; disagreements were resolved by consensus.

112 **Sample size and statistical analysis**

113 With 102 consecutively enrolled patients and an expected MACE incidence of approximately 20% in the LVEF-
114 preserved cohort, the study had 80% power at two-sided $\alpha = 0.05$ to detect an odds ratio of ≥ 3.5 for isolated DD
115 as a binary predictor—an effect size in keeping with prior estimates [14,24]. We acknowledge that the study was
116 underpowered for detecting smaller effects, and we therefore framed isolated-DD analyses as exploratory.

117 Categorical variables are reported as n (%); continuous variables as mean \pm standard deviation (SD) or median
118 (Q1, Q3) depending on distribution, assessed by the Shapiro–Wilk test. Two-group comparisons used the chi-
119 squared test or Fisher's exact test for categorical variables, and the Mann–Whitney U test for continuous
120 variables. Univariable and multivariable logistic regression identified MACE predictors; variable selection for
121 the multivariable model was guided by clinical prespecification rather than data-driven stepwise procedures, to
122 limit overfitting given the limited number of events per variable. Discriminative performance was assessed by
123 the area under the receiver-operating-characteristic curve (AUC) with bootstrap 95% CI (2,000 replicates). The
124 incremental value of DD added to a Lee + NT-proBNP base model was tested by paired-bootstrap AUC
125 comparison [25]. Internal validation used 1,000 bootstrap replicates to estimate optimism. Sensitivity,
126 specificity, and predictive values were calculated for isolated DD and elevated LVFP as standalone tests. All
127 analyses were performed with Python version 3.12 (SciPy version 1.17, scikit-learn version 1.8); two-sided P
128 values < 0.05 were considered significant. Reporting of the prediction-model component followed the TRIPOD
129 statement [26]. Missing data were minimal ($< 3\%$ per variable); complete-case analysis was used and no
130 imputation was performed.

131 **Declaration of use of generative artificial intelligence**

132 A large language model was used solely for language refinement and structural revision of the manuscript draft.
133 It was not used for study design, data collection, statistical analysis, or generation of figures or references. The
134 authors verified and take full responsibility for all content.

135 **Results:-**

136 **Cohort characteristics and echocardiographic phenotypes**

137 Of 102 analyzable patients (mean age 68.8 ± 10.3 years; 43.1% male), the echocardiographic phenotype
138 distribution was: Normal 36 (35.3%), Isolated DD 44 (43.1%), SD + DD 20 (19.6%), and Isolated SD 2 (2.0%)
139 (Fig. 1A). Among the 80 LVEF-preserved patients (78.4%), 44 (55.0%) met criteria for isolated DD—
140 numerically dominating the LVEF-preserved subgroup.

141 NT-proBNP differed strikingly across phenotypes (Fig. 1B): median 98 (Q1–Q3: 59 to 239) pg/ml in Normal
142 patients, 733 (173 to 981) pg/ml in Isolated DD, and 950 (414 to 3,216) pg/ml in SD + DD ($P < 0.001$ for
143 between-group comparison). Isolated DD patients therefore had a 7-fold higher median NT-proBNP than
144 phenotypically normal patients despite preserved LVEF—a biochemical signature of subclinical myocardial
145 stress.

146 **Clinical profile of isolated DD versus normal phenotype (LVEF-preserved cohort, n = 80)**

147 Isolated DD patients carried a substantially heavier cardiovascular comorbidity burden than Normal-phenotype
148 patients (Table 1; Fig. 2A): diabetes 59.1% vs. 16.7% ($P < 0.001$), hypertension 77.3% vs. 50.0% ($P = 0.021$),
149 prior cardiovascular history 72.7% vs. 44.4% ($P = 0.019$), NYHA ≥ 2 dyspnea 100% vs. 88.2% ($P = 0.051$,
150 trend), elevated LVFP 22.7% vs. 0% ($P = 0.002$), and NT-proBNP > 300 pg/ml in 63.6% vs. 22.2% ($P = 0.001$).

151 Importantly, age, sex, obesity, dyslipidemia, and smoking did not differ between groups, and the median Lee
152 score was identical (0 [0 to 0] vs. 0 [0 to 1], $P = 0.340$). Surgical risk class was also similar (high-risk surgery
153 22.7% vs. 16.7%, $P = 0.690$). This profile is consistent with isolated DD acting as a distinct pathophysiological
154 signature rather than a mere proxy for a heavier conventional risk panel (Table 1).

155 **Impact on preoperative management**

156 Detection of isolated DD significantly altered preoperative therapeutic decisions (Table 1; Fig. 2B):
157 preoperative therapeutic adjustments occurred in 50.0% of Isolated DD patients versus 11.1% of Normal
158 patients ($P < 0.001$; crude OR 8.00, 95% CI 3.00 to 21.00). Preoperative admission for optimization (40.9% vs.
159 22.2%, $P = 0.130$) and unplanned ICU admission (22.7% vs. 16.7%, $P = 0.690$) did not reach statistical
160 significance. The 5-fold higher rate of therapeutic adjustment reflects clinicians' real-world risk perception—
161 initiation or up-titration of beta-blockers, diuretics, statins, antithrombotics, and blood-pressure optimization—
162 and may itself have attenuated downstream MACE differences (an outcome paradox we address in the
163 Discussion).

164 **Independent prognostic value of isolated DD for MACE**

165 Sixteen of the 80 LVEF-preserved patients (20.0%) experienced perioperative MACE; rates were comparable
166 between phenotypes (Normal 16.7%, Isolated DD 22.7%, $P = 0.690$) (Table 1). On univariable analysis, isolated
167 DD was not a significant MACE predictor (OR 1.47, 95% CI 0.48 to 4.53; $P = 0.502$). Severe DD (isolated DD
168 plus elevated LVFP) trended toward a stronger effect (OR 3.22, 95% CI 0.79 to 13.19; $P = 0.104$), although the
169 analysis was power-limited (Table 2; Fig. 4).

170 The dominant univariable predictors of MACE were high-risk surgery (OR 16.11, 95% CI 4.32 to 60.05; $P <$
171 0.001), Lee score per point (OR 3.07, 95% CI 1.35 to 6.95; $P = 0.007$), and diabetes (OR 3.18, 95% CI 1.02 to
172 9.91; $P = 0.046$) (Table 2).

173 **Combined-model performance and incremental value of DD**

174 Predictive performance of competing models is shown in Table 4 and Fig. 3. Standalone discrimination of
175 isolated DD for MACE was poor (AUC 0.55; sensitivity 62%, specificity 47%, positive predictive value [PPV]
176 23%, negative predictive value [NPV] 83%). The Lee score alone yielded an AUC of 0.66; the Lee + NT-
177 proBNP base model reached 0.70.

178 Adding binary isolated DD to the Lee + NT-proBNP model produced no meaningful gain (AUC 0.70 \rightarrow 0.70;
179 Δ AUC = +0.002; bootstrap $P = 0.940$). In sharp contrast, adding severe DD (elevated LVFP) raised the AUC to
180 0.77—a +0.07 absolute improvement, consistent with a clinically relevant signal (Fig. 3B). The full
181 multivariable model integrating Lee + NT-proBNP + isolated DD + surgical risk achieved an apparent AUC of
182 0.89 (bootstrap-corrected 0.85). High-risk surgery ($\beta = +3.585$; OR 36.06, 95% CI 5.83 to 222.97; $P < 0.001$)
183 and Lee score ($\beta = +1.621$; OR 5.06, 95% CI 1.48 to 17.34; $P = 0.010$) retained significance; isolated DD alone
184 did not ($\beta = -0.910$; OR 0.40, $P = 0.394$) (Table 3).

185 **Diagnostic performance of isolated DD as a rule-out test**

186 As a standalone diagnostic test for MACE, isolated DD had a sensitivity of 62%, a specificity of 47%, a PPV of
187 23%, and an NPV of 83% (Table 5). Although the PPV is too low to drive escalation, the 83% NPV gives the
188 absence of DD a useful—if imperfect—rule-out role in routine preoperative practice.

189 **Discussion:-**

190 This prospective cohort study yields four findings that together motivate a reframing of preoperative diastolic
191 assessment in non-cardiac surgery. First, isolated DD with preserved LVEF affects more than half of LVEF-
192 preserved patients (55%)—a much higher prevalence than implied by current guideline texts [11,12]. Second, it
193 carries a robust biochemical and clinical signature: a 7-fold higher median NT-proBNP and a marked excess of
194 diabetes, hypertension, and prior cardiovascular disease, consistent with subclinical HFpEF [1,2]. Third, its
195 independent prognostic contribution to in-hospital MACE is, in fact, modest: as a binary predictor, it adds no
196 incremental discrimination over Lee + NT-proBNP. Fourth, and decisively, severity—captured by elevated
197 LVFP—does carry a clinically meaningful prognostic signal, lifting model AUC from 0.70 to 0.77.

198 Our findings sit coherently within the most recent evidence base. The 2024 AHA/ACC/ACS perioperative
199 guideline [11] explicitly notes that abnormal left ventricular systolic or diastolic function is associated with
200 increased perioperative MACE, but that the incremental value of NT-proBNP beyond risk indices and self-
201 reported functional capacity is small [18]. Our results extend this view: when DD is treated as binary, the
202 incremental signal disappears almost completely, mirroring the discrimination ceiling reported for binary
203 biomarker dichotomies. The Fayad et al. [14] systematic review and meta-analysis reported a pooled OR of
204 approximately 1.7 for any DD and MACE—an effect that is real but small. Higashi et al. [27] showed that an
205 $E/e' \geq 15$ (i.e., a severity criterion) tripled the risk of postoperative heart failure in intermediate-risk surgery,
206 again consistent with severity—not presence—as the operative dimension.

207 In the broader cardiology literature, DD severity has emerged as the dominant prognostic vector. The 2023 ESC
208 focused update on heart failure [20] elevated sodium–glucose cotransporter-2 (SGLT2) inhibitors (dapagliflozin
209 or empagliflozin) to a Class IA recommendation across the entire LVEF spectrum, including HFpEF, on the
210 basis of the EMPEROR-Preserved and DELIVER trials [32,33]. Recent network meta-analyses confirm
211 convergent benefit of SGLT2 inhibitors on cardiovascular death and heart-failure hospitalization in mildly
212 reduced and preserved ejection fraction [28]. Although these therapies are not deployed perioperatively, they
213 reframe the natural history we are trying to interrupt: preoperative isolated DD is not just an echocardiographic
214 finding but a prognostic entry point into a treatable trajectory.

215 Recent imaging work also resonates with our central message. The SOLOMON study [19] showed that
216 preoperative left ventricular global longitudinal strain modestly improved MACE prediction in non-cardiac
217 surgery patients—consistent with our finding that more granular indices of myocardial function add value where
218 binary classifications saturate. Meta-analyses of left atrial reservoir strain [29] support its prognostic role in
219 HFpEF and across surgical cohorts, suggesting a future axis for refining the severity gradient.

220 On the basis of our data and the broader literature, we propose a pragmatic preoperative algorithm for non-
221 cardiac surgery, which we term Severity-First Diastolic Triage (SFDT) (Fig. 5):

- 222 • Tier 1—Anchor on validated clinical predictors. Compute the Lee/RCRI score and identify high-risk
223 surgery as defined by 2022 ESC criteria [12]. These two variables retain by far the strongest independent
224 association with MACE in our and others' data.
- 225 • Tier 2—Reserve preoperative TTE for selected patients. Restrict routine preoperative echocardiography to
226 patients with NT-proBNP > 300 pg/ml, unexplained dyspnea, or known/suspected heart failure—as
227 already implied by the 2024 AHA/ACC [11] and 2022 ESC/ESAIC [12] guidelines, which discourage
228 indiscriminate preoperative TTE.
- 229 • Tier 3—Stratify by DD severity, not its binary presence. When TTE is performed, classify diastolic
230 function not only as present/absent but along the LVFP gradient. Isolated DD without elevated LVFP
231 confers a small absolute MACE excess that does not justify systematic escalation; isolated DD with
232 elevated LVFP markedly elevates risk and warrants preoperative cardiovascular optimization, anesthetic
233 plan adaptation (cautious volume management, avoidance of tachycardia, appropriate intraoperative
234 monitoring), and intensified postoperative surveillance.
- 235 • Tier 4—Use the absence of DD as a soft rule-out. The 83% negative predictive value of isolated DD alone
236 (combined with Lee = 0 and NT-proBNP < 300 pg/ml) supports straightforward progression to surgery
237 without further cardiac testing in the lowest-risk LVEF-preserved patients—aligning with the parsimony
238 favored by the 2024 AHA/ACC update [11].

239 This framework moves the conversation from “is there DD?”—a binary that drives both over-triage and under-
240 triage—to “how severe is the DD, and what does it mean alongside Lee and NT-proBNP?”. The full proposed
241 algorithm is summarized in Fig. 5.

242 Our 5-fold higher rate of preoperative therapeutic adjustments in isolated-DD patients (50.0% vs. 11.1%)
243 reveals an outcome paradox: clinicians already act on DD even when its independent prognostic value is small,
244 and that pre-emptive action may itself attenuate the MACE difference observed at the bedside [30]. This is not
245 an argument against assessment—it is an argument for targeted assessment, lest a finding that triggers expensive
246 escalation in many patients dilute its own observable risk effect through pre-emptive correction. The SFDT
247 framework formalizes this insight.

248 Pathophysiologically, the strong association of isolated DD with diabetes, hypertension, and elevated NT-
249 proBNP places these patients on the HFpEF continuum [1,2,31]. The preoperative consultation thus becomes an
250 unrecognized case-finding opportunity: identification of severe DD before surgery flags a population in whom
251 postoperative initiation of guideline-directed HFpEF therapy (SGLT2 inhibitors per the 2023 ESC update [20])
252 may have benefits well beyond the immediate perioperative window—a hypothesis worth testing in
253 interventional trials.

254 Strengths include the prospective design, consecutive enrollment, blinded echocardiographic assessment by a
255 single experienced operator using current 2016 ASE/EACVI criteria, complete preoperative biomarker capture,
256 and adjudicated outcomes. Reporting follows the STROBE and TRIPOD statements.

257 Several limitations deserve frank acknowledgment. First, the modest sample size (n = 80 in the primary
258 comparison) and the limited number of MACE events (n = 16) leave the study underpowered to detect small-to-

259 moderate independent effects of binary isolated DD. The non-significance of isolated DD as a MACE predictor
260 ($P = 0.502$) should be interpreted as absence of evidence, not evidence of absence. Second, single-center
261 recruitment limits external generalizability, particularly across surgical case-mix and ethnic backgrounds. Third,
262 the in-hospital composite endpoint mixes events of heterogeneous severity; longer-term outcomes (30-day, 90-
263 day, and 1-year MACE and mortality) were not captured. Fourth, although the operator was blinded, single-
264 reader echocardiographic interpretation introduces residual variability; multi-reader designs incorporating global
265 longitudinal strain and left atrial strain would strengthen future iterations. Fifth, high-sensitivity cardiac
266 troponin—now Class I in the 2022 ESC/ESAIC and 2024 AHA/ACC guidelines [11,12]—was not
267 systematically measured; its joint use with NT-proBNP and LVFP merits prospective study.

268 The SFDT framework requires external validation. Four research priorities follow:

- 269 • A multicenter prospective validation cohort (≥ 500 patients) powered to confirm the incremental
270 prognostic value of LVFP—not binary DD—over Lee + NT-proBNP, with longer follow-up (30-day and
271 1-year MACE).
- 272 • Integration of advanced echocardiographic markers—global longitudinal strain and left atrial reservoir
273 strain [19,29]—as continuous severity indices, potentially replacing the categorical LVFP threshold.
- 274 • A randomized pragmatic trial of “targeted TTE plus protocolized optimization” versus standard care in
275 NT-proBNP-positive patients, with hard cardiovascular outcomes.
- 276 • Implementation studies of SFDT-guided preoperative pathways with health-economic outcomes (cost per
277 MACE prevented, cardiology consultation rate, surgical-cancellation rate).

278 **Conclusion:-**

279 More than half of LVEF-preserved patients undergoing elective non-cardiac surgery have isolated diastolic
280 dysfunction. As a binary echocardiographic finding, isolated DD adds little to the prognostic discrimination
281 already achieved by the Lee score and NT-proBNP. Its severity—operationalized as elevated left ventricular
282 filling pressures—does carry a clinically meaningful incremental signal. Preoperative diastolic assessment
283 should therefore move from a binary alarm to a graded triage that integrates the Lee score, NT-proBNP, and
284 LVFP. The Severity-First Diastolic Triage framework operationalizes this shift and offers a pragmatic,
285 guideline-aligned pathway to rationalize preoperative decision-making in non-cardiac surgery.

286 **Conflicts of interest:-**

287 No potential conflict of interest relevant to this article was reported.

288 **Funding:-**

289 This research did not receive any specific grant from funding agencies in the public, commercial, or not-for-
290 profit sectors.

291 **Ethics approval and consent to participate:-**

292 The study protocol was approved by the institutional ethics committee (Comité d'Ethique pour la Recherche
293 Biomédicale, Université Mohammed V, Rabat; approval number 31/25) and complied with the Declaration of
294 Helsinki (revised 2013). Written informed consent was obtained from every participant prior to inclusion.

295 **Data availability:-**

296 The anonymized individual participant data and the analytic code that support the findings of this study are
297 available from the corresponding author upon reasonable request, subject to approval by the local Institutional
298 Review Board.

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- 415

416 **Tables**417 **Table 1.** Clinical characteristics and perioperative outcomes by diastolic status (LVEF-preserved cohort, n =
418 80).

419

Variable	Normal (n = 36)	Isolated DD (n = 44)	P value
Age (yr), mean \pm SD	67.6 \pm 11.9	69.5 \pm 9.4	0.900
Male sex, n (%)	10 (27.8)	16 (36.4)	0.570
Diabetes, n (%)	6 (16.7)	26 (59.1)	< 0.001
Hypertension, n (%)	18 (50.0)	34 (77.3)	0.021
Obesity, n (%)	8 (22.2)	14 (31.8)	0.480
Dyslipidemia, n (%)	8 (22.2)	8 (18.2)	0.870
Smoking, n (%)	4 (11.1)	8 (18.2)	0.530
Cardiovascular history, n (%)	16 (44.4)	32 (72.7)	0.019
Lee score, median (Q1, Q3)	0 (0, 0)	0 (0, 1)	0.340
Lee score \geq 1, n (%)	8 (22.2)	14 (31.8)	0.480
NT-proBNP (pg/ml), median (Q1, Q3)	98 (59, 239)	733 (173, 981)	< 0.001
NT-proBNP > 300 pg/ml, n (%)	8 (22.2)	28 (63.6)	0.001
NYHA \geq 2 dyspnea, n (%)	30/34 (88.2)	36/36 (100.0)	0.051
Elevated LVFP, n (%)	0 (0.0)	10 (22.7)	0.002
High-risk surgery, n (%)	6 (16.7)	10 (22.7)	0.690
Composite MACE, n (%)	6 (16.7)	10 (22.7)	0.690
Cardio-respiratory complications, n (%)	2 (5.6)	4 (9.1)	0.690
Postoperative ICU admission, n (%)	6 (16.7)	10 (22.7)	0.690
Therapeutic adjustment, n (%)	4 (11.1)	22 (50.0)	< 0.001
Preoperative admission, n (%)	8 (22.2)	18 (40.9)	0.130

420 *DD: diastolic dysfunction; ICU: intensive care unit; LVEF: left ventricular ejection fraction; LVFP: left ventricular filling pressures;*
 421 *MACE: major adverse cardiovascular events; NYHA: New York Heart Association; Q1, Q3: first and third quartiles; SD: standard*
 422 *deviation. Isolated DD was defined as preserved LVEF (\geq 50%) with abnormal diastolic function. Comparisons used the chi-squared or*
 423 *Fisher's exact test for categorical variables and the Mann-Whitney U test for continuous variables.*

424 **Table 2.** Univariable predictors of perioperative MACE (LVEF-preserved cohort, n = 80; 16 events).

425

Variable	OR (95% CI)	P value
Age (per 10 yr)	0.65 (0.38 to 1.11)	0.116
Diabetes	3.18 (1.02 to 9.91)	0.046
Hypertension	1.80 (0.52 to 6.22)	0.353
Obesity	1.80 (0.56 to 5.74)	0.320
Lee score (per point)	3.07 (1.35 to 6.95)	0.007
NT-proBNP > 300 pg/ml	2.44 (0.79 to 7.53)	0.122
High-risk surgery	16.11 (4.32 to 60.05)	< 0.001
Isolated DD (yes/no)	1.47 (0.48 to 4.53)	0.502
Severe DD (DD + elevated LVFP)	3.22 (0.79 to 13.19)	0.104
Elevated LVFP alone	3.22 (0.79 to 13.19)	0.104

426 *CI: confidence interval; DD: diastolic dysfunction; LVFP: left ventricular filling pressures; MACE: major adverse cardiovascular events;*
 427 *OR: odds ratio. Univariable logistic regression. Isolated DD as a binary predictor is non-significant; severity (elevated LVFP) trends*
 428 *toward a stronger association but is power-limited.*

429 **Table 3.** Multivariable logistic regression model of perioperative MACE.

430

Variable	β coefficient	OR (95% CI)	P value
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Variable	β coefficient	OR (95% CI)	P value
Constant (intercept)	-3.521	—	< 0.001
Lee score (per point)	+1.621	5.06 (1.48 to 17.34)	0.010
NT-proBNP > 300 pg/ml	+1.274	3.58 (0.47 to 27.14)	0.218
Isolated DD (yes/no)	-0.910	0.40 (0.05 to 3.26)	0.394
High-risk surgery	+3.585	36.06 (5.83 to 222.97)	< 0.001

431 *CI: confidence interval; DD: diastolic dysfunction; MACE: major adverse cardiovascular events; OR: odds ratio. Apparent model AUC =*
 432 *0.89; bootstrap-corrected AUC = 0.85 (1,000 replicates).*

433 **Table 4.** Discriminative performance of predictive models for perioperative MACE (area under the curve).

434

Model	AUC	Δ AUC vs. Lee + NT-proBNP
Lee score alone	0.66	—
NT-proBNP > 300 alone	0.61	—
Isolated DD alone	0.55	—
Elevated LVFP alone	0.58	—
Lee + NT-proBNP > 300	0.70	Reference
Lee + Isolated DD	0.67	—
Lee + NT-proBNP + Isolated DD	0.70	+0.002 (NS)
Lee + NT-proBNP + Severe DD (LVFP \uparrow)	0.77	+0.07
Full model (Lee + NT-proBNP + DD + Surg.)	0.89 / corrected 0.85	—

435 *AUC: area under the receiver-operating-characteristic curve; DD: diastolic dysfunction; LVFP: left ventricular filling pressures; MACE:*
 436 *major adverse cardiovascular events; NS: not significant. Binary DD does not improve the Lee + NT-proBNP base model; severity*
 437 *(elevated LVFP) provides a clinically meaningful gain (+0.07).*

438 **Table 5.** Diagnostic performance of isolated DD as a standalone MACE predictor (LVEF-preserved cohort, n =
 439 80).

440

Metric	Value (%)	Clinical interpretation
Sensitivity	62	62% of MACE occur in isolated-DD patients
Specificity	47	Low — many false positives
Positive predictive value (PPV)	23	23% of isolated-DD patients develop MACE
Negative predictive value (NPV)	83	83% of patients without isolated DD remain MACE-free

441 *DD: diastolic dysfunction; MACE: major adverse cardiovascular events; NPV: negative predictive value; PPV: positive predictive value.*
 442 *The 83% NPV gives the absence of isolated DD a useful—if imperfect—rule-out role in routine preoperative practice, supporting the rule-*
 443 *out tier of the Severity-First Diastolic Triage framework.*

444

445 **Figure legends:-**

446 **Fig. 1. Echocardiographic phenotypes and NT-proBNP distribution.** (A) Phenotype distribution in the
447 cohort (n = 102): Normal 35.3%, Isolated DD 43.1%, SD + DD 19.6%, Isolated SD 2.0%. (B) NT-proBNP by
448 phenotype (log scale); the median was 7-fold higher in Isolated DD than in Normal (733 vs. 98 pg/ml, P <
449 0.001). The dashed line indicates the 300 pg/ml ESC/ESAIC threshold. DD: diastolic dysfunction; ESAIC:
450 European Society of Anaesthesiology and Intensive Care; ESC: European Society of Cardiology; SD: systolic
451 dysfunction.

452 **Fig. 2. Clinical profile and perioperative outcomes—Isolated DD versus Normal phenotype (n = 80).** (A)
453 Comorbidities significantly more frequent in Isolated DD: diabetes, hypertension, cardiovascular (CV) history,
454 NYHA ≥ 2, elevated LVFP, and NT-proBNP > 300 pg/ml. (B) Perioperative outcomes: the composite MACE
455 did not differ, but a 5-fold higher rate of therapeutic adjustment occurred in Isolated DD (50% vs. 11%, P <
456 0.001). *** P < 0.001; ** P < 0.010; * P < 0.050; † P < 0.100. DD: diastolic dysfunction; ICU: intensive care
457 unit; LVFP: left ventricular filling pressures; MACE: major adverse cardiovascular events; NYHA: New York
458 Heart Association.

459 **Fig. 3. Receiver-operating-characteristic curves for perioperative MACE prediction.** (A) Standalone
460 predictors: Lee score (AUC 0.66), NT-proBNP > 300 (AUC 0.61), isolated DD (AUC 0.55), elevated LVFP
461 (AUC 0.58). (B) Combined models: Lee alone (AUC 0.66), Lee + NT-proBNP (AUC 0.70), and Lee + NT-
462 proBNP + Severe DD (AUC 0.77). The shaded area represents the AUC gain (+0.07) from adding DD
463 severity—not binary DD—to the base model. AUC: area under the receiver-operating-characteristic curve; DD:
464 diastolic dysfunction; LVFP: left ventricular filling pressures; MACE: major adverse cardiovascular events.

465 **Fig. 4. Forest plot of univariable predictors of perioperative MACE (n = 80; 16 events).** Odds ratios with
466 95% confidence intervals from univariable logistic regression. Red bars indicate P < 0.050; gray bars indicate P
467 ≥ 0.050. High-risk surgery (OR 16.11), Lee score (OR 3.07 per point), and diabetes (OR 3.18) are significant.
468 Binary isolated DD (OR 1.47) is not, whereas severity (DD + elevated LVFP) trends stronger (OR 3.22). The
469 dashed vertical line indicates OR = 1. CI: confidence interval; DD: diastolic dysfunction; LVFP: left ventricular
470 filling pressures; MACE: major adverse cardiovascular events; OR: odds ratio.

471 **Fig. 5. Severity-First Diastolic Triage (SFDT) framework—proposed preoperative pathway.** A four-tier
472 algorithm: (1) anchor on the Lee/RCRI score and the ESC surgical risk classification; (2) reserve preoperative
473 transthoracic echocardiography (TTE) for NT-proBNP > 300 pg/ml, unexplained dyspnea, or known/suspected
474 heart failure (HF); (3) stratify diastolic findings by the LVFP gradient, not binary DD; and (4) use absence of
475 DD as a soft rule-out. The framework is aligned with the 2024 AHA/ACC, 2022 ESC/ESAIC, and 2023 ESC
476 HF focused-update guidelines. AHA/ACC: American Heart Association/American College of Cardiology; DD:
477 diastolic dysfunction; ESC/ESAIC: European Society of Cardiology/European Society of Anaesthesiology and
478 Intensive Care; HF: heart failure; LVFP: left ventricular filling pressures; RCRI: Revised Cardiac Risk Index;
479 TTE: transthoracic echocardiography.

480

481 **Figures:-**

482

(A) Echocardiographic phenotype distribution (n = 102)

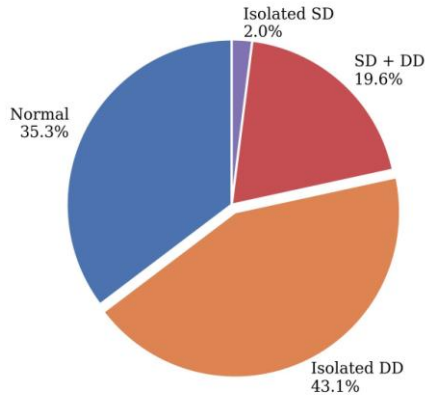
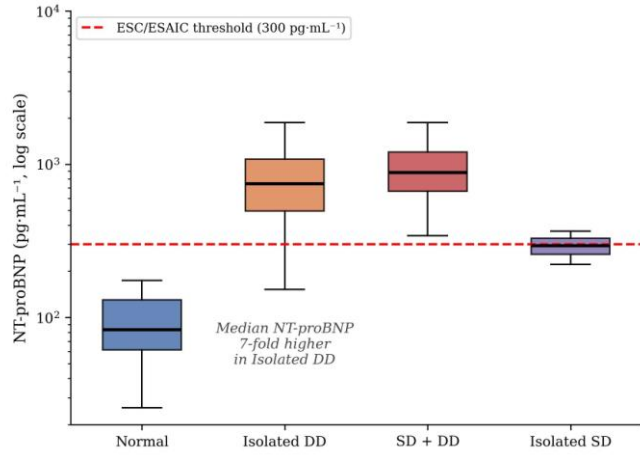


Fig. 1.

(B) NT-proBNP by phenotype (P < 0.001)

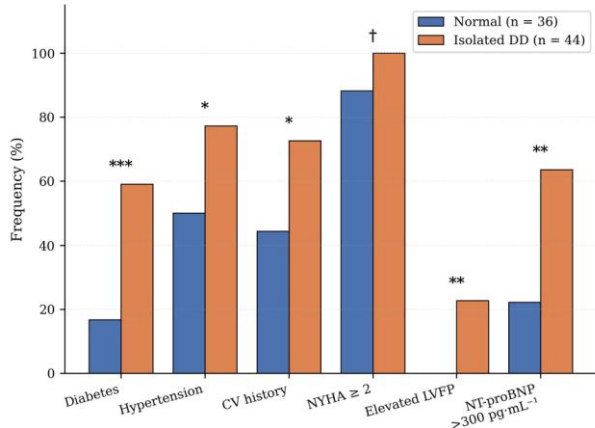


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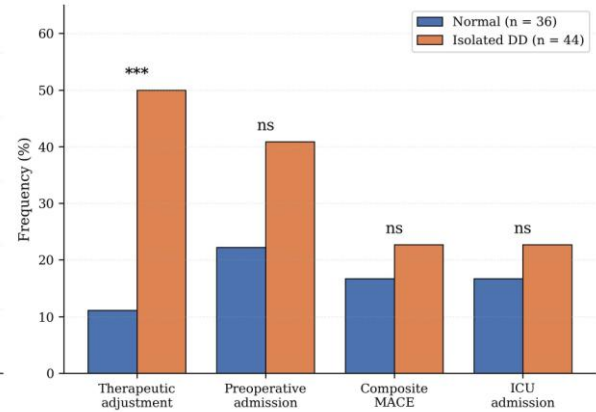
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Fig. 2.

(A) Cardiovascular comorbidities and biomarkers



(B) Perioperative outcomes



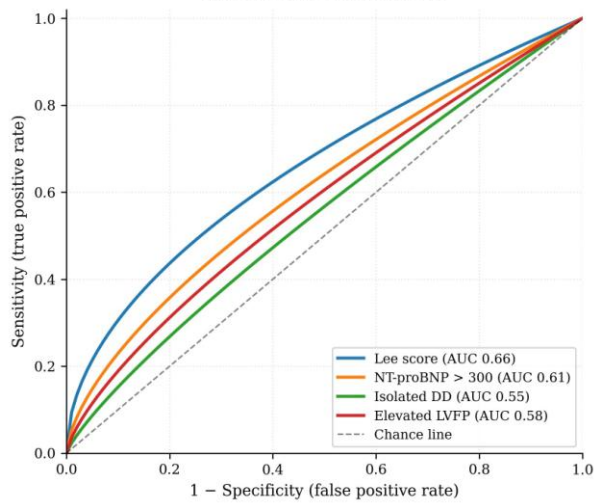
*** P < 0.001 ** P < 0.01 * P < 0.05 † P < 0.10 ns: not significant

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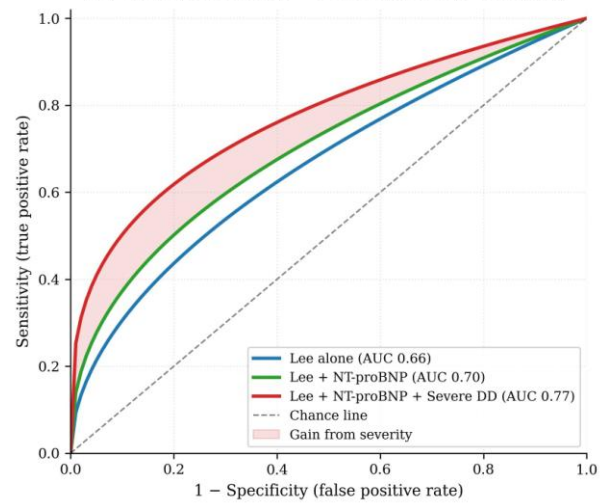
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Fig. 3.

(A) Standalone predictors



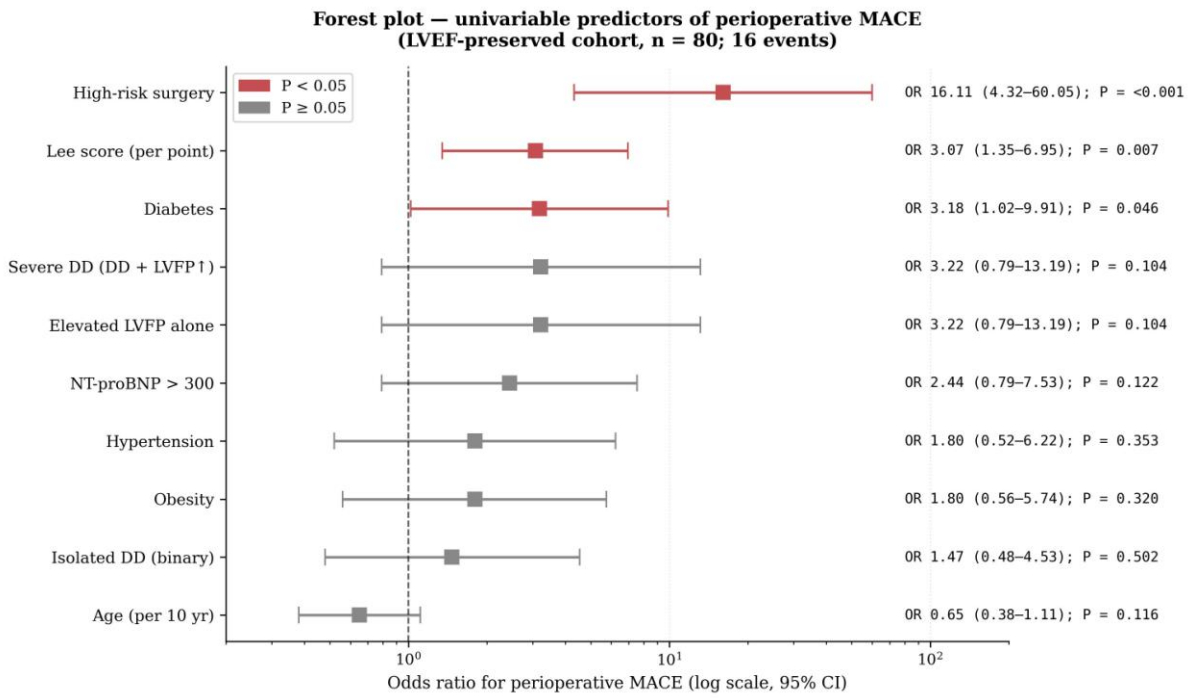
(B) Combined models — added value of DD severity



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Fig. 4.



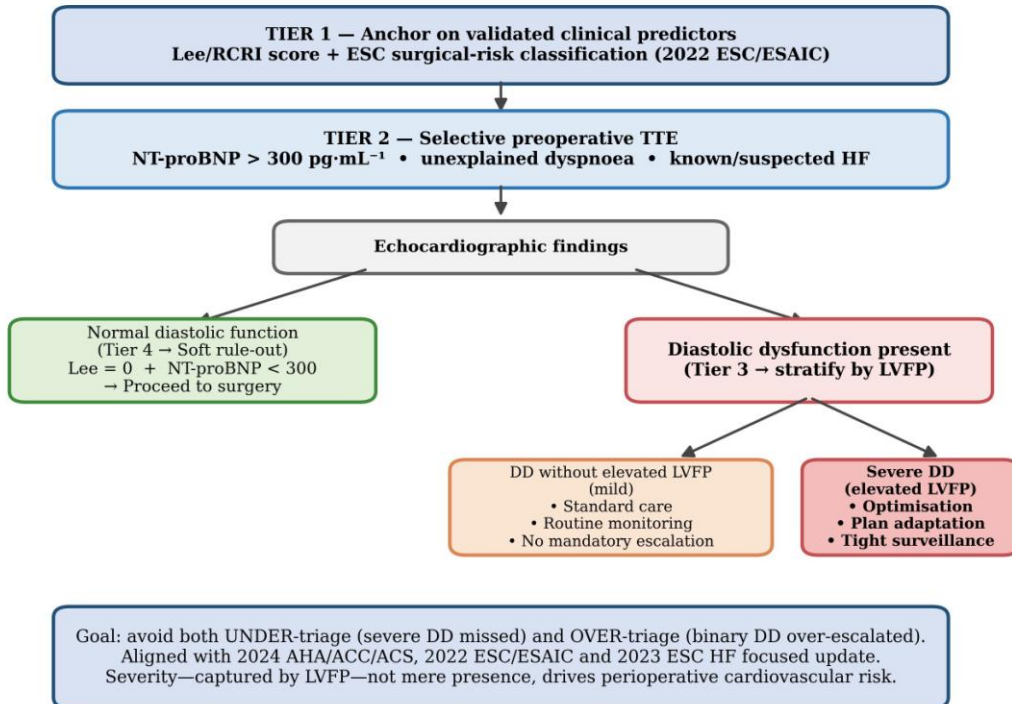
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Fig. 5.

Severity-First Diastolic Triage (SFDT) Framework

A pragmatic preoperative pathway for non-cardiac surgery



491