

Research Article

Using the Multidimensional Prognostic Index to Predict Clinical Outcomes of Hospitalized Older Persons: A Prospective, Multicenter, International Study

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See Acknowledgements for members of MPI_AGE Investigators

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Abstract

Background: Multidimensional Prognostic Index (MPI) is useful as a prognostic tool in hospitalized older patients, but our knowledge is derived from retrospective studies. We therefore aimed to evaluate in a multicenter, longitudinal, cohort study whether the MPI at hospital admission is useful to identify groups with different mortality risk and whether MPI at discharge may predict institutionalization, rehospitalization, and use of home care services during 12 months.

Methods: This longitudinal study, carried out between February 2015 and August 2017, included nine public hospitals in Europe and Australia. A standardized comprehensive geriatric assessment including information on functional, nutritional, cognitive status, risk of pressure sores, comorbidities, medications, and cohabitation status was used to calculate the MPI and to categorize participants in low, moderate, and severe risk of mortality. Data regarding mortality, institutionalization, rehospitalization, and use of home care services were recorded through administrative information.

Results: Altogether, 1,140 hospitalized patients (mean age 84.1 years, women = 60.8%) were included. In the multivariable analysis, compared to patients with low risk group at admission, patients in moderate (odds ratio [OR] = 3.32; 95% CI: 1.79–6.17; *p* < .001) and severe risk (OR = 10.72, 95% CI: 5.70–20.18, *p* < .0001) groups were at higher risk of overall mortality. Among the 984 older patients with follow-up data available, those in the severe-risk group experienced a higher risk of overall mortality, institutionalization, rehospitalization, and access to home care services.

Conclusions: In this cohort of hospitalized older adults, higher MPI values are associated with higher mortality and other negative outcomes. Multidimensional assessment of older people admitted to hospital may facilitate appropriate clinical and postdischarge management.

Keywords: Multidimensional Prognostic Index; Prognosis; Hospital; Cohort study; Aged

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Introduction

Increasing attention has been paid to the correct identification of mortality prognostic tools to help clinical decision making in diagnostics and therapeutics and to tailor appropriate interventions for the older frail patient (1).

Older adults are characterized by heterogeneous health status and health and functional trajectories, especially in the last years of life (2). Accordingly, clinical practice recommendations increasingly suggest that clinical decisions regarding older patients, including, for example, cancer screening or glycemic control in diabetes mellitus treatment, should take into account patients' life expectancy and/or prognosis for negative health outcomes (3,4).

The prognosis of complex older patients is not only ruled by the severity of individual diseases, but strongly related to the presence of multimorbidity and to the degree of physical, cognitive, biological, and social impairment (5). The comprehensive geriatric assessment (CGA), capable to effectively explore these domains, is the main tool to determine the prognosis of frail older persons (6). A systematic review of 16 common prognostic tools in geriatric medicine, however, identified very few prognostic indices for mortality that meet the requirements of accuracy and calibration required to be used in different settings (7).

Among the indices used in the hospital setting, the Multidimensional Prognostic Index (MPI) (8) has been identified as a well-calibrated tool with a good discrimination and accuracy both for short and long-term mortality (9). Moreover, the MPI is the only one based on information obtained from a CGA that explores comprehensively not only health aspects, but also functional, cognitive, and nutritional domains, as well as cohabitation status, using standardized and extensively validated rating scales, widely known by the clinicians (8). The MPI significantly predicts mortality in older patients hospitalized for different reasons, but data to support this come mainly from retrospective studies (8–14).

The aim of this study was to prospectively evaluate whether the CGA-based MPI at admission to hospital is useful to identify different mortality risk groups in terms of increased survival during 12 months of follow-up in a multicenter and longitudinal cohort study. Moreover, we investigated whether MPI at discharge is useful to predict other outcomes significantly associated with higher risk of mortality, such as institutionalization, rehospitalization, and use of home care services.

Methods

Study Population

This was an observational study conducted according to the World Medical Association's 2008 Declaration of Helsinki, the guidelines for Good Clinical Practice, and the Strengthening the Reporting of Observational Studies in Epidemiology (STROBE) guidelines (15).

Inclusion criteria were the following: (a) age more than 65 years, (b) admitted to hospital for an acute medical condition or for a relapse of a chronic disease, (c) ability to provide an informed consent or availability of a proxy for informed consent and willingness to participate in the study, and (d) complete CGA during hospitalization. Nine geriatric units across Europe and Australia were included. The recruitment period was from February 2015 to July 2016, with the end of 1-year follow-up in August 2017.

The ethical committees of each center approved this observational study. Informed consent was given by participants who underwent initial evaluation and/or their proxies for their clinical records to be used in this study. All patient records and information were anonymized and de-identified prior to the analysis.

Main Exposure: MPI

MPI was developed by the inclusion of information from eight different domains of the CGA (8):

- Functional status was evaluated by Katz's Activities of Daily Living (ADL) index (16), which defines the level of dependence/ independence in six daily personal care activities (bathing, toileting, feeding, dressing, urine and bowel continence, and transferring) in and out of bed or chair.
- 2. Independence in the Lawton's Instrumental Activities of Daily Living (IADL) (17), which assesses independence in eight activities that are more cognitively and physically demanding than ADL, that is, managing finances, using telephone, taking medications, hopping, using transportation, preparing meals, doing housework, and washing.
- 3. Cognitive status through the Short Portable Mental Status Questionnaire (SPMSQ) (18), a 10-item questionnaire investigating orientation, memory, attention, calculation, and language; validated versions were used in each local language.
- 4. Comorbidity was examined using the Cumulative Illness Rating Scale (CIRS) (19). The CIRS uses a 5-point ordinal scale (score 1–5) to estimate the severity of pathology in each of 13 systems, including cardiac, vascular, respiratory, eye-ear-nose-throat, upper and lower gastrointestinal, hepatic, renal, genitourinary, musculo-skeletal, skin disorder, nervous system, endocrino-metabolic, and psychiatric behavioral disorders. On the basis of the ratings, the Comorbidity Index (CIRS-CI) score, which reflects the number of concomitant diseases, was derived from the total number of categories in which moderate or severe levels (grade from 3 to 5) of disease were identified (range from 0 to 13). Comorbidities, at hospital admission, were descriptively reported using the *International Classification of Disease, Tenth Revision* (20).
- Nutritional status was investigated with the Mini Nutritional Assessment–Short Form (MNA-SF) (9), which includes information on: (a) anthropometric measures (body mass index, weight loss), (b) neuropsychological problems and recent psychological stress, (c) mobility, and (d) decline in food intake.
- 6. Risk of developing pressure sores was evaluated through the Exton Smith Scale (ESS), a five-item questionnaire determining physical and mental condition, activity, mobility, and incontinence (21).
- 7. Medication use was defined according to the Anatomical Therapeutics Chemical Classification code system (ATC classification) and the number of drugs used by patients at admission was recorded. Patients were defined as drug users if they took a medication included in the ATC classification at the moment of hospital admission.
- 8. Cohabitation status included living alone, in an institution, or with family members.

For each domain, a tripartite hierarchy was used, that is, 0 = no problems, 0.5 = minor problems, and 1 = major problems, based on conventional cutoff points derived from the literature for the singular items. The sum of the calculated scores from the eight domains was divided by 8 to obtain a final MPI risk score ranging from 0 = no risk to 1 = higher risk of mortality. Also, the MPI was expressed as three grades of risk: MPI-1, low risk (MPI value ≤ 0.33); MPI-2, moderate risk (MPI value between 0.34 and 0.66); and MPI-3, high

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risk (MPI value > 0.66) (8). MPI requires between 15 and 25 minutes for its complete execution and the results can be automatically obtained through the program Calculate-MPI, which can be download for free by the www.mpiage.eu website. The program is an.exe file and allows to run out the tests to obtain the numeric value of the single domain (ADL, IADL, SPMSQ, CIRS, MNA, ESS, drugs, cohabitation) as well as the final MPI score. Tests are available also in printable PDF format, as shown in the Supplementary Material.

Main Outcomes

Participants were followed for 12 months from hospital discharge and mortality was categorized as in-hospital (if it happened in the index hospital admission) or postdischarge. Vital status was assessed, when appropriate, by consulting the Registry Offices of the cities in which the patients were residents at the time of the evaluation.

Overall mortality (as sum of in-hospital and 1 year, at home) was considered as primary outcome of our research. Institutionalization, rehospitalization, and access to home care services were considered as secondary end points. These outcomes were investigated using administrative data and phone interviews with patients and/or caregivers.

Statistical Analysis

A sample size of 1,000 participants was deemed sufficient, assuming a fixed Type I error alpha of 5% and a statistical power of 80%, to detect an increase in mortality at 1 year equivalent to a risk of at least 36% for those participants classified as MPI-2 and at least 37% for those participants classified as MPI-3, set the MPI class 1 as the reference. One-year mortality was taken for the reference class equal to 10% (22).

General characteristics were reported as frequencies (percentages) and mean \pm standard deviation, for categorical and continuous variables, respectively. Mortality incidence rates were computed as the number of deaths per person-year. Comparisons across MPI categories were performed using the Kruskal–Wallis test for singly ordered contingency tables, and linear by linear association test, for categorical and continuous variables, respectively.

Logistic binary regression analysis was run, taking MPI at admission (in categories or as increase in 0.1 points) as exposure and overall (in-hospital + postdischarge) mortality, in-hospital mortality, and postdischarge (1-year) follow-up mortality as outcomes. The strength of the association between MPI at admission and these outcomes were reported as odds ratios (ORs) with 95% confidence intervals (CIs), adjusted for age, gender, center, and primary diagnosis. We also reported the adjusted risk that come from the adjusted predicted probability of the outcome of interest for each level of MPI, when the other covariates in the model are at their means. For all the outcomes, receiver operator characteristic curves were analyzed to compare the sensitivity and specificity of MPI in predicting

Table 1.	General	Characteristics	of Older	Patients	at Baseline	According to I	MPI Grade
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		All Patients	MPI-1 Mild Risk	MPI-2 Moderate Risk	MPI-3 Severe Risk	
Variable Mean (SD)		1,140	169 (14.8%)	502 (44.0%)	469 (41.2%)	p*
Length of stay (mean,	SD)	14.0 (11.8)	8.0 (5.6)	13.6 (10.5)	16.6 (13.6)	<.001
Age (mean, SD)		84.2 (7.4)	81.5 (6.9)	83.8 (7.2)	85.6 (7.4)	<.001
Gender $(n, \%)$	Female	694 (60.9)	84 (49.7)	319 (63.5)	291 (62.0)	.005
	Male	446 (39.1)	85 (50.3)	183 (36.5)	178 (38.0)	
ADL		2.8 (2.3)	5.6 (0.7)	3.6 (1.9)	0.8(1.1)	<.001
IADL		2.7 (2.6)	6.4 (1.7)	3.1 (2.2)	0.8 (1.1)	<.001
SPMSQ		4.1 (3.5)	1.2 (1.4)	2.7 (2.5)	6.7 (3.1)	<.001
MNA-SF		8.6 (3.3)	11.4 (2.1)	9.7 (2.6)	6.4 (2.9)	<.001
ESS		14.1 (3.7)	18.3 (1.6)	15.6 (2.5)	11.0 (2.8)	<.001
CIRS		2.1 (0.4)	1.8 (0.4)	2.1 (0.4)	2.3 (0.4)	<.001
Number of medication	15	7.1 (3.4)	5.2 (3.1)	7.3 (3.6)	7.6 (2.9)	<.001
Cohabitation status	Alone	372 (32.7)	46 (27.4)	191 (38.0)	135 (28.8)	<.001
(<i>n</i> , %)	with family or private care	623 (54.6)	122 (71.4)	269 (51.8)	232 (47.8)	
	home care/in institution	144 (12.6)	0 (0.0)	42 (8.4)	102 (21.7)	
Main diagnosis* by	AB: Infectious	87(7.8)	4 (2.4)	33 (6.6)	50(11.1)	<.001
ICD-10	CD: Neoplasms	55 (4.9)	7 (4.2)	29 (5.8)	19 (4.2)	.51
(<i>n</i> , %):	D5: Blood	34 (3.1)	10 (6.0)	20 (4.0)	4 (0.9)	<.001
	E: Endocrine, nutritional and metabolism	41 (3.7)	12 (7.1)	16 (3.2)	13 (2.9)	.05
	FG: Nervous system and behavioral	81 (7.3)	10 (6.0)	32 (6.4)	39 (8.7)	.36
	HLT: Skin, ear, and others	25 (2.2)	4 (2.4)	8 (1.6)	13 (2.9)	.42
	I: Circulatory system	260 (23.3)	46 (27.4)	123 (24.8)	91 (20.2)	.10
	J: Respiratory system	174 (15.6)	25 (14.9)	71 (14.3)	78 (17.3)	.43
	K: Digestive system	103 (9.2)	19 (11.3)	45 (9.1)	39 (8.7)	.57
	M: Musculoskeletal and connective	73 (6.5)	16 (9.5)	28 (5.6)	29 (6.4)	.21
	N: Genitourinary system	36 (3.2)	4 (2.4)	16 (3.2)	16 (3.6)	.80
	RS: Fractures, tendency to fall, edema	147 (13.2)	11 (6.5)	76 (15.3)	60 (13.3)	.01

Notes: ADL = Activities of Daily Living; CIRS = Comorbidity Index Rating Scale; ESS = Exton-Smith Scale; IADL = Instrumental Activities of Daily Living; ICD-10 = International Classification Disease, Tenth Revision; MNA-SF = Mini Nutritional Assessment–Short Form; SD = standard deviation; SPMSQ = Short Portable Mental Status Questionnaire

*Diagnosis was missing in 24 patients.

these negative outcomes, measuring the relative area under the curve with the correspondent 95% CI.

After excluding people dead during hospitalization and those with no data at follow-up, in secondary analyses, MPI at discharge was used for analyses having overall mortality, institutionalization, rehospitalization, and access to home care services as outcomes.

Two-sided alternatives with a significance level (alpha = 0.05) were considered for all the tests. STATA 12 (Stata Statistical Software: Release 12; StataCorp, College Station, TX) software was used.

Results

Characteristics of the Study Population

The study population included 1,140 hospitalized patients (mean age 84.2 ± 7.4 years, women = 60.8%).

Table 1 shows the characteristics of participants according to their MPI grade at admission: 169 patients (14.8%) were in the MPI-1 (mild risk of mortality group), 502 patients (44.0%) in the MPI-2 (moderate-risk group), and 469 patients (41.2%) in the MPI-3 (severe risk). Participants with higher MPI values (ie, MPI-3, severe risk) were more likely to be females (p = .005) and older (p < .001) as well as of having significantly worse scores in all the domains considered in the MPI, as expected (p < .001 for all comparisons). Participants classified in the MPI-3 group (high risk) lived more frequently in institutions (p < .001) and had a longer length of stay in hospital (p < .001). Regarding main diagnoses at hospital discharge, participants included in the higher MPI-risk group had a significant higher prevalence of infectious diseases (p < .001) or fractures/tendency to fall (p = .01) than participant included in the MPI-1 low-risk group.

MPI at Admission and Mortality

Table 2 shows the association between MPI (at admission) and mortality. In the multivariate analysis, taking participants in the lowrisk group as reference, those in the moderate (OR = 3.32, 95% CI: 1.79-6.17, p < .001) and high-risk groups (OR = 10.72, 95% CI: 5.70–20.18, p < .001) were at significantly higher risk of mortality. Similar findings were evident when taking postdischarge (1-year) and in-hospital mortality as outcomes.

Modelling MPI at admission as continuous variable, each increase in 1 decimal point in MPI at admission was significantly associated with a higher risk of overall (OR = 1.60, 95% CI: 1.45–1.76, p < 1.45.001), in-hospital mortality (OR = 1.54, 95% CI: 1.29-1.86, p < .001), and postdischarge (1-year) mortality (OR = 1.57, 95% CI: 1.42–1.74, *p* < .001) (Table 2).

Finally, MPI had a good discriminatory power for all the end points considered, as showed by the correspondent values of area under the curves reported in Table 2 being 0.77 (95% CI: 0.74–0.80; Figure 1a) for overall mortality, 0.79 (95% CI: 0.73–0.85; Figure 1b) for in-hospital mortality, and 0.76 (95% CI: 0.73-0.79; Figure 1c) for 1-year mortality.

Prognostic Value of MPI at Hospital Discharge

In 984 older participants, the MPI was available also at discharge. During hospitalization, MPI improved (ie, delta MPI < 0) in 350 participants (35.6%), worsened in 264 participants (ie, delta MPI > 0; 26.8%) whereas MPI did not change in 370 participants (ie, delta MPI = 0; 37.6%). As reported in Table 3, taking those with MPI-1 mild risk as reference and after adjusting the analyses for potential confounders, participants in the severe-risk group at discharge experienced a higher risk of mortality (OR = 7.36, 95% CI: 3.87–13.97,

			MPI at Admission				
Outcome	Events/Total (%)	Estimates	Mild Risk	Moderate Risk	Severe Risk	Increase in 0.1 Point	AUC ⁺ (95% CI)
Overall (in-hospital + post discharge) mortality	358/1,140 (31.4)	Adjusted risk (95% CI)	8.0% (3.7–12.2)	22.4% (18.3–26.5)	48.2% (42.7–53.7)	OR = 1.60 (1.45-1.76) $h < .001$	0.77 (0.74–0.80)
		Odds ratio, OR (95% CI) <i>p</i> -value. OR	reference	3.32 (1.79–6.17) b < 001	10.72 (5.70-20.18) b < 0.01		
In-hospital mortality	75/1,140 (6.6)	Adjusted risk (95% CI) Odds ratio, OR (95% CI)	0.7% (0.0–1.7) reference	2.5% (1.2–3.9) OR = 3.78 (0.81–17.66)	6.4% (3.9-9.0) OR = 9.99 (2.18-45.80)	OR = 1.54 (1.29–1.86)	0.79 (0.73–0.85)
Postdischarge (1-year) follow-up mortality	283/1,004‡(31.4)	<i>p</i> -value, OR Adjusted risk (95% CI) Odds ratio, OR (95% CI) <i>p</i> -value, OR	reference 6.9% (3.0–10.9) reference reference	p = .09 19.1% (15.2 - 23.0) OR = 3.17 (1.64-6.13) $p = .001$	p = .003 42.8% (37.1-48.5) OR = 10.04 (5.12-19.68) $p < .001$	P < .001 OR = 1.57 (1.42-1.74) p < .001	0.76 (0.73–0.79)
Notes: All data (except for AUCs) :	are reported as odds ratic	os (ORs) are adjusted for age, ger	nder, center, and primar	ry diagnosis. AUC = area under i	the curve; CI = confidence interva	ıls; MPI = Multidin	nensional Prognostic

Table 2. Logistic Regression on Baseline MPI Influencing Mortality Risk in Hospitalized Older Patients

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^{‡75} participants were dead during hospitalization and 61 were lost at the follow-up [†]AUCs were related to model with MPI in category.



Figure 1. Receiver operating characteristic (ROC) curves for baseline Multidimensional Prognostic Index (MPI) in predicting overall (a), in-hospital (b), and 1-year (c) mortality.

p < .001), institutionalization (OR = 2.15, 95% CI: 1.07–4.32, p = .03), rehospitalization (OR = 1.62, 95% CI: 1.02–2.56, p = .04), and access to home care services (OR = 1.89, 95% CI: 1.01–3.52, p = .046) (Table 3). Using the MPI at discharge as continuous variable did not significantly change our findings (Table 3).

Similarly to MPI at admission, MPI at discharge had a good discriminatory power for all the end points considered, as showed by the correspondent values of area under the curves ranging from 0.65 (95% CI: 0.61–0.69) for rehospitalization to 0.81 (0.79–0.85) for institutionalization (Table 3).

Discussion

In this real-world prospective observational study of a large cohort of hospitalized older individuals in different countries with different health care systems, we found that MPI was a significant predictor not only of mortality (both short and long term), but also of other negative outcomes, such as institutionalization, rehospitalization, and higher use of home care services. MPI showed a good discriminatory power and accuracy in predicting all the end points included in our investigation.

Because of the expansion in number of old and very old patients in the population, a progressively larger percentage of the hospitalized patients are old and frail. The development of prognostic tools appropriate for the evaluation of hospitalized older patients is capturing increasing attention. Recent guidelines suggest to incorporate life expectancy as a key factor in weighing the benefits and the burdens of both diagnostic and therapeutic interventions (23-26). Prognostic indices may offer an important tool to the clinician for better understanding the appropriate clinical decision making for older adults (5). It is widely known that failure to consider prognosis in the context of clinical decision making can lead to poor care (4). For example, hospice is often underutilized for patients with nonmalignant yet life-threatening diseases (27), whereas healthy older patients with good prognosis have low rates of cancer screening precluding them early interventions (7,28). Thus, simple and reliable tools for the estimation of the prognosis of the older patients are needed to tailor clinical management of older patients.

The MPI has been validated in more than 12,000 older patients with acute or relapses of chronic conditions (such as pneumonia, heart failure, gastrointestinal bleeding, chronic renal failure, liver cirrhosis, dementia, and transient ischemic attack, using mortality as outcome (22,29). A previous multicenter study, involving more than 2,000 hospitalized older patients, has shown that MPI is a more accurate prognostic predictor than three frailty indices commonly used in clinical practice for mortality (30). Even if of importance, these data were derived from already collected retrospective data (8–14). The research reported here was specifically designed and conducted

			MPI at Discharge				
Dutcome	Events/Total (%)	Estimates	Mild Risk	Moderate Risk	Severe Risk	Increase in 0.1 Point	AUC [†] (95% CI)
Ostdischarge (1-year) ollow-up mortality	283/1,004‡(28.2)	Adjusted risk (95% CI) Odds ratio, OR (95% CI) <i>b</i> -value. OR	10.2% (4.9–15.4) reference reference	17.3% (13.5-21.2) OR = 1.85 (0.98-3.49) b = .06	45.4% (39.3-51.6) OR = 7.36 (3.87-13.97) b < .001	OR = 1.57 (1.42- 1.75) b < .001	0.78(0.74–0.81)
nstitutionalization	$189/1,004^{\ddagger}(18.8)$	Adjusted risk (95% CI) Odds ratio, OR (95% CI) <i>b</i> -value. OR	7.9% (3.2–12.5) reference reference	$\begin{array}{l} \begin{array}{c} r \\ r \\ 13.4\% (9.6-17.3) \\ OR = 1.82 (0.95-3.51) \\ p = .07 \end{array}$	15.5% (10.7-20.2) OR = 2.15 (1.07-4.32) b = .03	OR = 1.19 (1.06-1.33) b = .003	0.81 (0.78–0.85)
Rehospitalization	<i>572/</i> 1,004 [‡] (<i>57.</i> 0)	Adjusted risk (95% CI) Odds ratio, OR (95% CI) <i>p</i> -value. OR	45.3% (36.1–54.5) reference reference	$p_{\rm e} = 0.7\% (55.8 - 65.6)$ 0R = 1.87 (1.21 - 2.88) p = .004	57.2% (51.4-63.1) OR = 1.62 (1.02-2.56) b = .04	OR = 1.07 OR = 1.07 (0.99-1.16) p = .07	0.65 (0.61–0.69)
Access to home care services	s 251/1,004 [#] (25.0)	Adjusted risk (95% CI) Odds ratio, OR (95% CI) <i>p</i> -value, OR	15.7% (8.3–23.1) reference reference	$\sum_{j=1}^{2} 5.3\% (20.2-30.3)$ OR = 1.81 (1.01-3.26) p = .048	26.0% (20.2–31.9) OR = 1.89 (1.01–3.52) <i>p</i> = .046	$p = -\frac{1}{2}$ OR = 1.05 (0.95-1.16) p = -31	0.78 (0.74–0.81)
Notes: All data (except :	for AUCs) are reported	as odds ratios (ORs) are adjusted	for age, gender, center, and	primary diagnosis. AUC = area u	nder the curve; CI = confidence in	tervals; MPI = Multidii	mensional Prognostic

Table 3. Logistic Regression on MPI at Discharge and Negative Outcomes in Hospitalized Older Patients

⁴⁷⁵ participants were dead during hospitalization and 61 were lost at the follow-up

[†]AUCs were related to model with MPI in category.

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to confirm the retrospective findings mentioned before. We confirmed that the MPI is significantly associated with a higher risk of mortality in older hospitalized patients. Compared to the previous works regarding MPI, the prospective design can overcome the intrinsic limitations of retrospective studies, in particular recall bias of the participants, the absence of a priori hypothesis and, sometimes, the presence of missing data not recorded at the baseline (31).

Another finding of our research is that the MPI score confirms to be sensitive to variations of health status during hospitalization. Indeed, in almost 1,000 hospitalized participants with data at discharge, more than one third had a substantial improvement in their MPI score, whereas in about one fourth we observed a worsening. These findings confirm previous results that MPI is able to capture the modifications of the patient clinical conditions over hospital stay (32) and suggest that it could be used to monitor the effectiveness of interventions in hospitalized older patients.

This work for the first time reports that MPI is a significant predictor not only of mortality, but also of other negative outcomes of public health importance in older people, such as rate of institutionalization, rehospitalization, and use of home care services. These findings might be of importance for clinicians and health care managers in order to identify not only people at higher mortality risk, but also those with specific needs of health or social care. Although one third of the patients included in our analyses died during the 1 year of follow-up, more than 50% were rehospitalized and about one quarter were placed in nursing home or needed home-care services. As our study found that the risk of these negative outcomes significantly increases with the increase of MPI score, our findings suggest that MPI could be used for identifying people with potentially higher care needs. Very interesting, MPI accurately predicts the negative health outcomes independently from the different health and social care organizations of the participant centers and countries. As these results were based on a large multicenter study, without selective exclusion criteria, with different causes of admission to the hospital and consequently including patients observed in daily practice, we also believe that MPI could be considered as having a good external validity and so applied for better following patients after their discharge at home.

The findings of our study should be interpreted within its limitations. First, the follow-up of these patients was limited to 1 year; therefore, we cannot exclude that significant differences in predicting negative outcomes by MPI in older patients with different mortality risk could emerge with longer follow-up. Moreover, we did not include the competing risk by death for the analysis of the secondary outcomes, possibly introducing a bias. Second, even if the power of MPI in detecting negative outcomes may be considered good (1,5), the area under the curves were lower than 0.90, thus suggesting unexplained variation in mortality. However, it should be noted that other indices that commonly drive clinical decisions, such as the congestive heart failure, hypertension, age >75 years, diabetes mellitus, prior stroke 2 or transient ischemic attak, or thromboembolism (CHADS2) index to help determine warfarin therapy (33), the Framingham risk score to help determine lipid therapy (34), and the thrombolysis in myocardial infarction risk score to help determine invasive therapy for unstable angina (35), had similar values in C-index. Finally, even if most of the scales included in the MPI are well known and routinely collected by health care professionals involved in many geriatric programs, it is possible that in other settings the execution of the MPI can require a specific training and more time than that reported in the present study.

In conclusion, in this large cohort of hospitalized older adults, a significant association between higher MPI values and higher allcause mortality and other negative outcomes was found, over 1 year of follow-up. These findings clearly indicate the value of assessing older people admitted to hospital in order to choose the most appropriate clinical management for these individuals. Therefore, future randomized controlled trials with different kinds of multidimensional interventions and including a consistent part of people having different degrees of frailty are needed to confirm or not the present observations.

Supplementary Material

Supplementary data is available at *The Journals of Gerontology, Series A: Biological Sciences and Medical Sciences* online.

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Conflict of Interest Statement

None declared.

References

- Siontis GC, Tzoulaki I, Ioannidis JP. Predicting death: an empirical evaluation of predictive tools for mortality. *Arch Intern Med.* 2011;171:1721– 1726. doi:10.1001/archinternmed.2011.334
- Gill TM, Gahbauer EA, Han L, Allore HG. Trajectories of disability in the last year of life. N Engl J Med. 2010;362:1173–1180. doi:10.1056/ NEJMoa0909087
- Schoenborn NL, Bowman TL II, Cayea D, Pollack CE, Feeser S, Boyd C. Primary care practitioners' views on incorporating long-term prognosis in the care of older adults. *JAMA Intern Med.* 2016;176:671–678. doi:10.1001/jamainternmed.2016.0670
- Lee SJ, Leipzig RM, Walter LC. Incorporating lag time to benefit into prevention decisions for older adults. *JAMA*. 2013;310:2609–2610. doi:10.1001/jama.2013.282612
- Gill TM. The central role of prognosis in clinical decision making. JAMA. 2012;307:199–200. doi:10.1001/jama.2011.1992
- Pilotto A, Cella A, Pilotto A, et al. Three decades of comprehensive geriatric assessment: evidence coming from different healthcare settings and specific clinical conditions. J Am Med Dir Assoc. 2017;18:192.e1–192. e11. doi:10.1016/j.jamda.2016.11.004
- Yourman LC, Lee SJ, Schonberg MA, Widera EW, Smith AK. Prognostic indices for older adults: a systematic review. JAMA. 2012;307:182–192. doi:10.1001/jama.2011.1966

- Pilotto A, Ferrucci L, Franceschi M, et al. Development and validation of a multidimensional prognostic index for one-year mortality from comprehensive geriatric assessment in hospitalized older patients. *Rejuvenation Res.* 2008;11:151–161. doi:10.1089/rej.2007.0569
- Sancarlo D, D'Onofrio G, Franceschi M, et al. Validation of a Modified-Multidimensional Prognostic Index (m-MPI) including the Mini Nutritional Assessment Short-Form (MNA-SF) for the prediction of one-year mortality in hospitalized elderly patients. J Nutr Health Aging. 2011;15:169–173.
- Bureau ML, Liuu E, Christiaens L, et al.; MPI_AGE Project Investigators. Using a multidimensional prognostic index (MPI) based on comprehensive geriatric assessment (CGA) to predict mortality in elderly undergoing transcatheter aortic valve implantation. *Int J Cardiol.* 2017;236:381–386. doi:10.1016/j.ijcard.2017.02.048
- Pilotto A, Addante F, Ferrucci L, et al. The multidimensional prognostic index predicts short- and long-term mortality in hospitalized geriatric patients with pneumonia. J Gerontol A Biol Sci Med Sci. 2009;64:880– 887. doi:10.1093/gerona/glp031
- Pilotto A, Addante F, Franceschi M, et al. Multidimensional Prognostic Index based on a comprehensive geriatric assessment predicts short-term mortality in older patients with heart failure. *Circ Heart Fail*. 2010;3:14– 20. doi:10.1161/CIRCHEARTFAILURE.109.865022
- Sancarlo D, Pilotto A, Panza F, et al. A Multidimensional Prognostic Index (MPI) based on a comprehensive geriatric assessment predicts short- and longterm all-cause mortality in older hospitalized patients with transient ischemic attack. J Neurol. 2012;259:670–678. doi:10.1007/s00415-011-6241-4
- 14. Volpato S, Bazzano S, Fontana A, Ferrucci L, Pilotto A; MPI-TriVeneto Study Group. Multidimensional Prognostic Index predicts mortality and length of stay during hospitalization in the older patients: a multicenter prospective study. J Gerontol A Biol Sci Med Sci. 2015;70:325–331. doi:10.1093/gerona/glu167
- von Elm E, Altman DG, Egger M, Pocock SJ, Gøtzsche PC, Vandenbroucke JP; STROBE Initiative. The Strengthening the Reporting of Observational Studies in Epidemiology (STROBE) statement: guidelines for reporting observational studies. J Clin Epidemiol. 2008;61:344–349. doi:10.1016/j. jclinepi.2007.11.008
- Katz S, Downs TD, Cash HR, Grotz RC. Progress in development of the index of ADL. *Gerontologist*. 1970;10:20–30.
- Lawton MP, Brody EM. Assessment of older people: self-maintaining and Instrumental Activities of Daily Living. *Gerontologist.* 1969;9:179–186. doi:10.1093/geront/9.3_Part_1.179
- Pfeiffer E. A Short Portable Mental Status Questionnaire for the assessment of organic brain deficit in elderly patients. J Am Geriatr Soc. 1975;23:433441. doi:10.1097/00006199-199911000-00009
- Linn BS, Linn MW, Gurel L. Cumulative illness rating scale. J Am Geriatr Soc. 1968;16:622–626. doi:10.1111/j.1532-5415.1968.tb02103.x
- Harris ST, Zeng X, Ross T, Ford L. International Classification of Diseases, 10th Revision training: what coders are saying. Health Care Manag (Frederick). 2014;33:91–93. doi:10.1097/HCM.0b013e3182a9d640
- Bliss MR, McLaren R, Exton-Smith AN. Mattresses for preventing pressure sores in geriatric patients. *Mon Bull Minist Health Public Health Lab Serv.* 1966;25:238–268.

- 22. Pilotto A, Panza F, Ferrucci L. A multidimensional prognostic index in common conditions leading to death in older patients. Arch Intern Med. 2012;172:594; discussion 594–595. doi:10.1001/ archinternmed.2011.1891
- 23. Brown AF, Mangione CM, Saliba D, Sarkisian CA; California Healthcare Foundation/American Geriatrics Society Panel on Improving Care for Elders with Diabetes. Guidelines for improving the care of the older person with diabetes mellitus. J Am Geriatr Soc. 2003;51 (suppl 5 Guidelines):S265–S280. doi:10.1046/j.1532-5415.51.5s.1.x
- 24. Force USPST. Colon cancer screening (USPSTF recommendation). U.S. Preventive Services Task Force. J Am Geriatr Soc. 2000;48:333–335.
- Mottet N, Bellmunt J, Bolla M, et al. EAU-ESTRO-SIOG guidelines on prostate cancer. Part 1: screening, diagnosis, and local treatment with curative intent. *Eur Urol.* 2017;71:618–629. doi:10.1016/j.eururo.2016.08.003
- Freedman RA, Keating NL, Pace LE, Lii J, McCarthy EP, Schonberg MA. Use of surveillance mammography among older breast cancer survivors by life expectancy. J Clin Oncol. 2017;35:3123–3130. doi:10.1200/ JCO.2016.72.1209
- 27. Casarett D, Karlawish J, Morales K, Crowley R, Mirsch T, Asch DA. Improving the use of hospice services in nursing homes: a randomized controlled trial. *JAMA*. 2005;294:211–217. doi:10.1001/ jama.294.2.211
- Walter LC, Covinsky KE. Cancer screening in elderly patients: a framework for individualized decision making. JAMA. 2001;285:2750–2756.
- 29. Pilotto A, Sancarlo D, Daragjati J, Panza F. Perspective: the challenge of clinical decision-making for drug treatment in older people. The role of multidimensional assessment and prognosis. *Front Med (Lausanne)*. 2014;1:61. doi:10.3389/fmed.2014.00061
- 30. Pilotto A, Rengo F, Marchionni N, et al.; FIRI-SIGG Study Group. Comparing the prognostic accuracy for all-cause mortality of frailty instruments: a multicentre 1-year follow-up in hospitalized older patients. *PLoS One.* 2012;7:e29090. doi:10.1371/journal.pone.0029090
- Vandenbroucke JP. Observational research, randomised trials, and two views of medical science. *PLoS Med.* 2008;5:e67. doi:10.1371/journal. pmed.0050067
- 32. Volpato S, Daragjati J, Simonato M, Fontana A, Ferrucci L, Pilotto A. Change in the multidimensional prognostic index score during hospitalization in older patients. *Rejuvenation Res.* 2016;19:244–251. doi:10.1089/ rej.2015.1715
- 33. Rietbrock S, Heeley E, Plumb J, van Staa T. Chronic atrial fibrillation: incidence, prevalence, and prediction of stroke using the Congestive heart failure, Hypertension, Age >75, Diabetes mellitus, and prior Stroke or transient ischemic attack (CHADS2) risk stratification scheme. Am Heart J. 2008;156:57–64. doi:10.1016/j.ahj.2008.03.010
- 34. D'Agostino RB Sr, Grundy S, Sullivan LM, Wilson P; CHD Risk Prediction Group. Validation of the Framingham coronary heart disease prediction scores: results of a multiple ethnic groups investigation. *JAMA*. 2001;286:180–187.
- 35. Antman EM, Cohen M, Bernink PJ, et al. The TIMI risk score for unstable angina/non-ST elevation MI: a method for prognostication and therapeutic decision making. *JAMA*. 2000;284:835–842.