

1Introduction

2Hospital readmissions represent an increasing, significant and highly expensive cost for National
3Health Services. A hospital readmission is defined as an admission to an acute care hospital within
430 days of discharge from the same or another acute care hospital.¹

5In the United States, preventable hospital readmissions were estimated to contribute over \$25
6billion to wasteful health care spending annually.² Previous research has shown that among the
7nine million hospitalizations of Medicare patients per year³, almost one fifth of patients are
8readmitted within a month of discharge and many more return to the emergency room.⁴

9Rehospitalization shortly after discharge is increasingly recognized as a marker of inpatient quality
10of care⁵, but has been observed that current hospital readmission measures are inadequate for
11reporting hospital quality. Readmissions are in fact influenced by multiple sources of variation,
12such as mix of patients served or bias in the performance measure, and true differences in quality
13of care are often a much smaller source of this variation.⁶

14Geographic variation is also closely related to hospital readmissions, depending on the chances to
15access the specific medical care for a disease⁷, and also disparate physician and family
16expectations may affect both the decision-making process and patients' care outcomes resulting in
17repeated hospitalizations.⁸

18Zapatero et al.⁹ showed that readmissions within 30 days to Spanish Internal Medicine
19departments are prevalent as 12.4%, and key factors that predicted subsequent admission
20included male sex, length of stay, comorbidities, and some clinical conditions. In Italy, the overall
21hospital readmission prevalence was 10.2%, with a total of 43.7% judged to be potentially
22avoidable.¹⁰

23In countries with a rapidly ageing population, increasing life expectancy, rising chronic disease
24burden and decreasing working population are all factors that contribute to straining the finite
25healthcare resource further.^{11,12} Higher comorbidity rate^{13,14} and multidrug resistant infections¹⁵
26have been shown to be associated with an increased risk of readmission.

27There is consensus that in order to reduce avoidable hospital readmissions, it is crucial to properly
28identify the patients at highest risk, and main modifiable risk factors¹⁶.

29Thus, assessing the role of comorbidities in determining repeated readmissions in real-life is
30crucial, but there is a need for more data. Additional research on understanding this complex
31phenomenon, and to find effective interventions applicable to acute inpatient populations aimed
32to reduce potentially avoidable readmissions, are strongly needed.

33Internal Medicine and Geriatric Unit,^{9,17} are notoriously very affected by this problem; therefore,
34they can provide updated real-life data to understand the evolution of the phenomenon over
35time.

36In this short paper, we retrospectively analyzed all patients who underwent readmissions to our
37Italian Internal Medicine ward during a year, assessing the main demographic characteristics of
38patients, their comorbidities and mortality, and performing an analysis depending by the DRGs
39distribution, comparing them with controls who underwent only one hospitalization episode.

40 41Materials and Methods

42A retrospective case-control study was conducted between January 1 and December 31, 2018 at
43the Internal Medicine Unit of the Hospital of Pontedera (Pisa, Italy), assessing the characteristics of
44all consecutive patients who required hospitalization. Among these, we compared the patients
45who underwent hospital readmissions (cases) with those patients who required only one
46hospitalization during the year and were discharged alive (controls).

47In particular, the main demographic characteristics of all patients, their comorbidities and
48mortality, the home-patient setting and social problems previously emerged, were carefully

49assessed. All the patients received a comprehensive assessment within 24 hours from admission,
50administered by trained clinicians.

51The 30-days re-admissions were retrospectively analyzed crossing the data collected from the
52Hospital Discharge Register containing the Diagnosis-Related Group (DRG) codes. All data were
53retrieved matching data from our electronic health record (Sistema C7, Fondazione Gabriele
54Monasterio, CNR Pisa) with our DRG software system (ADT, Engineering Group, Roma)¹⁸. Among
55all the readmissions, the most relevant and frequent DRG was included in the study. Moreover, we
56performed an analysis depending by the DRGs distribution among cases and controls.

57All DRGs were grouped into clusters, according to the main organ involvement or pathogenetic
58mechanism.¹⁹ Table 1S shows the DRG clustering and the average weight of each major diagnostic
59category observed in the study.

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61Statistical Analysis

62Continuous variables were defined as mean \pm standard deviation; categorical variables were given
63as percentage. The independent sample t test was used for the continuous variables and the chi-
64square test for categorical variables. A logistic regression analysis was also performed. Significance
65was inferred for a p value <0.05. Statistical analysis was performed using Microsoft Excel version
662016.

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68Results

69In the 12-months study period a total of 3012 patients (mean age 76.8 ± 14.3 years; range 17-102,
7046.7% males) were admitted to our Internal Medicine Unit.

71Cases, defined as patients who underwent hospital readmissions, were 426 (14.1%), their mean
72age was 79.7 ± 11.9 years (range 23-100), and 45.8% were males.

73Controls were 420 (13.9%), their mean age was 75.9 ± 14.7 years (range 22-99), 44.3% were males.

74Among the 426 cases, the number of readmissions ranged from a minimum of 2 to a maximum of
758 in the study period (Table 1).

76Table 2 shows the results of the univariate analysis. The mean age was higher in the case group
77(79.7 vs 75.9 years), while no significant difference was found between groups for the sex.

78Cases showed a higher prevalence of cerebrovascular disease (77.2% vs 48.1%), cognitive decline
79(51.8% vs 26.9%), congestive heart failure (47.6% vs 20.2%), chronic kidney disease (31.7% vs
8013.1%), chronic obstructive pulmonary disease (23.0% vs 14.5%), skin ulcers (45.1% vs 17.6%) than
81controls. No significant difference was found regarding diabetes and active cancer.

82We assessed also the presence of skin ulcers, showing a significantly higher proportion among
83patients requiring readmissions (45.1% vs 17.6%).

84Regarding patient setting, provenience from home-based health care settings was significantly
85higher in controls group (91.9% vs 84.7%), while previous extended care was significantly more
86frequent in patients who required readmissions (12.7% vs 6.2%). Previously identified social
87problems were instead similarly distributed between the two groups.

88The report of diagnosis related groups (DRGs) in both groups is listed in table 3. The average
89length of stay (ALOS) was similar (9.2 ± 7.1 days in cases vs 10.0 ± 7.0 days in controls), such as the
90DRG average weight.

91Main characteristics and DRGs of dead patients are reported in Table 4. DRGs analysis showed a
92higher proportion of Infectious disease (24.4% vs 15.0%) and Kidneys-Urinary Tract-Reproductive
93organs related diseases (9.6% vs 5.0%) in the case group than in controls. Conversely, Central
94Nervous System disease (22.4% vs 10.6%) and Gastrointestinal-liver related diseases (10.5% vs
956.6%) were significantly higher in the controls group. All other DRGs did not show significant
96changes in distribution between cases and controls.

97The absolute mortality rate was significantly higher in cases than in controls (53.5% vs 37.4%),
98such as the occurrence of death within 30-days from last admission (38.2% vs 12.1%). The average
99time between admission and death was significantly longer in controls group (57.7 versus 31.0
100days). The mean age of patients who died was similar in cases and controls.
101At the DRGs analysis, Sepsis (31.6% vs 19.1%), Pneumonia (17.1% vs 7.6%) and Kidney Failure
102(9.6% vs 3.8%) were significantly more frequently in cases compared to controls the main cause of
103mortality. For controls, a significantly higher proportion of Active Cancers (25.5% vs 17.1%) was
104observed. Among dead patients, DRG average weight was significantly higher in controls than in
105cases (1.20 vs 1.14)

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107**Discussion**

108In our study, cases who underwent hospital readmissions within 30 days were 14.1% of total of
 1093012 patients hospitalized during a year in an Internal Medicine ward, a percentage similar to that
 110of 12.4% reported by Zapatero et al.⁶ in Spanish Internal Medicine departments. Unlike these
 111authors, we have not found male sex and length of stay as significant key factors predicting
 112readmissions. A reduction of length of stay, especially in elderly subjects, increased early
 113rehospitalization rates also in the work of Fabbian et al.²⁰ conducted among patients admitted to
 114the department of medicine of a general hospital in Italy. In the Fabbian' study the most frequent
 115causes of rehospitalization were cardiovascular disease in 29.3% and pulmonary disease in 29.7%
 116of cases; also in our cohort, considering distribution among cases, Lung disease was the DRG most
 117frequently reported in 32.6%, while Cardiovascular disease was less often codified (4.5%).
 118Our study shows a high complexity of patients who underwent repeated hospital readmissions,
 119since they are older and very often affected by chronic diseases.

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120Considering the baseline characteristics of patients, we have not observed statistical differences
121for diabetes and active cancer. Regarding diabetes, this could be related to the high and increasing
122prevalence of this condition in general population for whom has been there for some time an
123extensive and structured health assistance network in the outpatient setting, while regarding
124active cancer an efficient model of management (the Home-based health care was even
125significantly more frequent in controls) and awareness of the poor prognosis associated with the
126disease, could explain why the number of readmissions is not higher in this category of very fragile
127patients. Thereafter, patients who underwent readmissions were significantly more affected by
128chronic diseases that can progress to a fast worsening or exacerbations, such as Chronic Kidney
129Disease and Chronic obstructive pulmonary disease.

130Cognitive decline was significantly more represented (twice fold) in patients who required
131readmissions, confirming what is already known from the current literature.²¹

132In this study, the presence of skin ulcers were specifically assessed as key factors determining the
133readmission risk, showing a significant impact as a major determinant. The chronic course of ulcers
134needs the creation of a structured territorial network capable of managing them, otherwise they
135lead to continuous re-hospitalization both directly and indirectly for local and systemic
136complications, such as infections and sepsis. The role of skin ulcers deserves more attention than
137what has been given so far: in a recent multinational study, the combinations of diseases
138categories that were most strongly associated with readmissions, were chronic kidney disease
139with liver disease or chronic skin ulcers.²²

140Regarding to patients setting provenience, we observed that the majority of patients in both
141groups were previously treated in a Home-based health care, with a weak higher significance for
142controls group. Patients who required readmissions, more often came from nursing homes or
143similar health-assisted settings (extended care), suggesting that the out-of-hospital system is
144evidently not yet completely ready to take charge of a large number of patients with chronic
145situations. A recent meta-analysis showed that at 1 month from discharge, the continuity of care
146interventions was associated with lower readmission rates, preventing short-term hospital
147readmission in older people with chronic diseases, but without any conclusive evidence about the
148effectiveness aiming to reduce long-term readmission.²³ Many episodes of readmission can be
149ultimately related to social environment, poor familiar assistance and inadequate follow-up
150program, anyway the suitability of early rehospitalization as a correct target for good medical
151practice is to date debated.²⁴

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152 Social problems, often related to poor economic conditions, were low and equally distributed in
153 both groups: in our study, they did not represent a relevant factor for recurrent hospitalizations.
154 Apart the lung disease-related DRG, representing the most represented in both groups, Infectious
155 disease was the first DRG in the readmission group (24.4%), significantly higher than controls
156 (15.0%). It is well known that readmissions after a sepsis episode are very common and costly, and
157 that the majority of readmissions were associated with infectious aetiologies, thus efforts are
158 needed to clarify all the determinants of readmission after a first sepsis episode and develop
159 strategies in terms of quality of care and care transitions to prevent this outcome.²⁵
160 Central nervous system DRGs were more frequent (twice) in controls group: a possible explanation
161 is that, being control patients relatively less affected by chronic comorbidities and being the
162 assignment of DRG mainly based on the highest resources consumption, these DRGs can emerge
163 even if they are less remunerative than those related to lung diseases and sepsis, for whom there
164 is a stronger relationship with the revolving group.
165 Absolute mortality rate (53.5%) was higher, as expected, compared with other data in literature
166 regarding patients with readmissions,²⁶ but once again, we have to underline that sepsis was our
167 first DRG in dead cases, followed by pneumonia and kidney failure: our data is in line with the high
168 mortality related to sepsis when occurring in elderly patients.²⁷
169 Cancer-related DRGs were instead significantly more frequent in dead controls: this is interesting,
170 suggesting that, since controls are less affected by chronic illnesses, their DRG really emerges as
171 cancer. Anyway, DRG average weight was then significantly higher in control group, since cancer,
172 and in particular haematological malignancies, have a highly remunerative DRG.
173 No significant differences were observed for the other death-related DRGs between groups.
174 We noted that the high percentage of revolving patients with any skin ulcer (45.1%) did not have
175 an adequate correspondence in DRG (Skin Wound 1.4%): this could be explained both by a
176 tendency to underestimate the importance of ulcers remuneration, and by the fact that skin ulcers
177 in such patients frequently represent the source of sepsis, which is subsequently encoded in the
178 main DRG.
179 We believe that focusing on the problem of ulcers could represent a strength of our study.
180 A main limitation of this study is that, for the retrospective design, only the association between
181 factors, not causation, can be determined. Moreover, since DRGs are operator-encoded
182 procedures, there could be some bias and thus a variability in the results, but the operator who
183 encoded DRG in our Department was a very expert and trained clinician.

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185 Conclusions

186 In conclusion, our study confirms that readmissions to Internal Medicine Departments are
187 frequent and mainly related to the severity of chronic diseases affecting the patients, especially
188 when able to sudden worsening towards an acute phase. Sepsis was the most relevant condition

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189associated with readmissions, leading to a very high mortality rate, and this could indirectly reflect
190the global burden of antibiotic resistance: Italy is currently one of the most interested country in
191Europe.²⁸ Skin ulcers are present in about half of the patients who are early readmitted, but they
192were coded as DRG only in an absolute minority, so more accurate coding is needed. Although
193further investigation is warranted in order to clarify these complex relationships, preventing sepsis
194and investing resources in chronic diseases assistance represent a key challenge for the future.

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