Since January 2020 Elsevier has created a COVID-19 resource centre with free information in English and Mandarin on the novel coronavirus COVID-19. The COVID-19 resource centre is hosted on Elsevier Connect, the company's public news and information website.

Elsevier hereby grants permission to make all its COVID-19-related research that is available on the COVID-19 resource centre - including this research content - immediately available in PubMed Central and other publicly funded repositories, such as the WHO COVID database with rights for unrestricted research re-use and analyses in any form or by any means with acknowledgement of the original source. These permissions are granted for free by Elsevier for as long as the COVID-19 resource centre remains active.
Original article

Malnutrition risk as a negative prognostic factor in COVID-19 patients

Mancin Stefano*a, Bertone Andreat, Cattani Daniela b, Morenghi Emanuelaa, b, Passadori Lorenaa, Donizetti Daniela a, Fanny Søkeland b, Elena Azzolini a, Mazzoleni Beatriceb
a IRCCS Humanitas Research Hospital Rozzano, Milan, Italy
b Humanitas University Pieve Emanuele, Milan, Italy

A R T I C L E   I N F O

Article history:
Received 16 July 2021
Accepted 20 July 2021

Keywords:
Malnutrition
SARS-CoV-2
Infection
Immune system
Obesity
ACE2

S U M M A R Y

Background/objective: SARS CoV-2 infection is a disease, whose prevalence has drastically risen in the past year. The aim of this study is to examine a possible association between the risk of malnutrition, clinical outcomes following hospitalisation and morbidity at discharge.

Methods: This study has analysed the medical records of 652 patients hospitalised at Humanitas Research Hospital (Milan, Italy) between 01/03 and 30/04/2020. The risk of malnutrition was identified with the Malnutrition Universal Screening Tool (MUST).

Results: The cohort was composed of 515 patients. The MUST scale is significantly associated to malnutrition evaluating the morbidity at discharge (discharged 0.27 ± 0.68, discharged with problems 0.40 ± 0.93, deceased 0.64 ± 0.93, p < 0.001), and the clinical outcome following hospitalisation (HR 1.25, 95% CI 1.04–1.51, p = 0.019) is maintained even after correction for age, treated hypertension, admission to an intensive care unit and oxygen therapy). A subgroup analysis addressing patients with a BMI ≥30 shows a significant association between comorbidities such as: arterial hypertension (HR 4.95, 95% CI 1.10–22.22, p = 0.037), diabetes (HR 3.37, 95% CI 1.04–10.89, p = 0.043) and renal failure (HR 3.94, 95% CI 1.36–11.36, p = 0.011).

Conclusions: The results of this study suggest that the risk of malnutrition is a noteworthy indicator that impacts both the clinical outcomes and morbidity at discharge.

© 2021 European Society for Clinical Nutrition and Metabolism. Published by Elsevier Ltd. All rights reserved.

1. Introduction

Infection from SARS-CoV-2 has shown to lead to a condition of extreme clinical severity. Since its first appearance until present no definitive cure has been found, even with ongoing massive research efforts. Many studies have identified risk factors for unfavourable outcome, preventive measures and treatments, to improve the prognostic factors of the disease. Malnutrition has been recognized as one of these factors [1].

Malnutrition is defined as a condition of functional, organic and developmental alteration resulting from an imbalance between required nutrients, actual intake and metabolism leading to an increase in mortality and morbidity affecting life quality. Such a clinical condition can result from reduced intake of macro- and micronutrients, excessive intake or from an altered nutrient metabolism [2].

Prevalence rates of malnutrition in hospitalised patients in Europe vary according to the criteria used for identification. On average 35% of newly admitted patients (range 10–80%) presents malnutrition, which is generally aggravated during hospitalisation [3].

Malnutrition is the main cause of immunodeficiency in the world, affecting both the innate and the acquired immune response [4], exposing individuals to an elevated risk of infection [5] and a lower capacity to inhibit viral proliferation.

The immune system can further be altered by malnutrition due to excess food intake. Obesity induces a systemic inflammatory state with consequential alteration TCD4 cell response brought about by leptin, increasing autoimmunity [6] along with T-cell exhaustion characterised by reduced effective, proliferative and cytotoxic function [7].

Micronutrient deficiencies are a growing problem in individuals with malnutrition, as vitamins and micronutrients play an important role in both immune and other metabolic systems. Micronutrient deficiencies are a major cause of immunosuppression, which may contribute to the increased risk of infection in both adult and paediatric patients [8].

1.1. Malnutrition

Malnutrition is a clinical condition that affects patients with a BMI < 21. It is defined as a condition of functional, organic, and developmental alteration resulting from an imbalance between required nutrients, actual intake, and metabolism, leading to an increase in mortality and morbidity affecting life quality. Such a clinical condition can result from reduced intake of macro- and micronutrients, excessive intake or from an altered nutrient metabolism [2].

Prevalence rates of malnutrition in hospitalised patients in Europe vary according to the criteria used for identification. On average 35% of newly admitted patients (range 10–80%) presents malnutrition, which is generally aggravated during hospitalisation [3].

Malnutrition is the main cause of immunodeficiency in the world, affecting both the innate and the acquired immune response [4], exposing individuals to an elevated risk of infection [5] and a lower capacity to inhibit viral proliferation.

The immune system can further be altered by malnutrition due to excess food intake. Obesity induces a systemic inflammatory state with consequential alteration TCD4 cell response brought about by leptin, increasing autoimmunity [6] along with T-cell exhaustion characterised by reduced effective, proliferative and cytotoxic function [7].

Micronutrient deficiencies are a growing problem in individuals with malnutrition, as vitamins and micronutrients play an important role in both immune and other metabolic systems. Micronutrient deficiencies are a major cause of immunosuppression, which may contribute to the increased risk of infection in both adult and paediatric patients [8].

1.2. Immunodeficiency

Immunodeficiency is a clinical condition that affects patients with a BMI < 21. It is defined as a condition of functional, organic, and developmental alteration resulting from an imbalance between required nutrients, actual intake, and metabolism, leading to an increase in mortality and morbidity affecting life quality. Such a clinical condition can result from reduced intake of macro- and micronutrients, excessive intake or from an altered nutrient metabolism [2].

Prevalence rates of malnutrition in hospitalised patients in Europe vary according to the criteria used for identification. On average 35% of newly admitted patients (range 10–80%) presents malnutrition, which is generally aggravated during hospitalisation [3].

Malnutrition is the main cause of immunodeficiency in the world, affecting both the innate and the acquired immune response [4], exposing individuals to an elevated risk of infection [5] and a lower capacity to inhibit viral proliferation.

The immune system can further be altered by malnutrition due to excess food intake. Obesity induces a systemic inflammatory state with consequential alteration TCD4 cell response brought about by leptin, increasing autoimmunity [6] along with T-cell exhaustion characterised by reduced effective, proliferative and cytotoxic function [7].

Micronutrient deficiencies are a growing problem in individuals with malnutrition, as vitamins and micronutrients play an important role in both immune and other metabolic systems. Micronutrient deficiencies are a major cause of immunosuppression, which may contribute to the increased risk of infection in both adult and paediatric patients [8].

© 2021 European Society for Clinical Nutrition and Metabolism. Published by Elsevier Ltd. All rights reserved.
role in allowing correct functioning of the innate and acquired im-

mune response [8–10]. A state of malnutrition associated with a
deficit in micronutrients, hypermetabolism and an excessive loss of
nitrogen, are all factors known to predispose infection [11].

Furthermore, obesity has a strong link to respiratory diseases. It
is widely known to be associated with asthma, obstructive sleep
apnoea, acute lung damage and ARDS [12] (acute respiratory
distress syndrome). Therefore, the association of severe pulmonary
complications from COVID-19 and obesity is now assessed, and
obesity is a known risk factor for severe outcome [13–16] also in
other form of pandemic influenza like H1N1 [17].

2. Materials and methods

A retrospective cohort study was carried out. The primary
objective was to evaluate the possible association between the risk
of malnutrition, evaluated at the moment of hospitalisation for
respiratory distress caused by a COVID-19 infection, and clinical
outcomes such as: mortality, length of stay and morbidity at
discharge. The presence of pressure sores, the occurrence of su-
perinfections and the admission to intensive care units were ana-
lysed as secondary outcomes.

Inclusion criteria were: adult patients admitted from the
Emergency Department with a diagnosis of respiratory distress
given by a SARS-CoV-2 infection. Exclusion criteria were: patients
who were transferred directly from the Emergency Department to
Intensive care units, patients for whom an initial assessment could
not be carried out and the patients who did not give consent for the
use of their personal data for research.

Data of all hospitalised patients admitted directly from the
Emergency Department to Humanitas Research Hospital, a uni-
versity tertiary hospital in Rozzano (Milan, Italy) wards between 1
March and 30 April 2020, when respecting the inclusion/exclusion
criteria, were collected retrospectively in May 2020, for a total of
652 patients. The data collected comprehend information about
patient general conditions at admission (vital signs, pre-existing
comorbidities and concomitant diseases) and nutritional state,
with the aim of assigning a value of nutritional risk on the
Malnutrition Universal Screening Tool (MUST) complemented with
an evaluation of the nutritional state of the patient.

2.1. Literature review and identification of a rating scale for
nutritional risk

A literature review was carried out in April 2020, using the main
databases. The criteria used for the review were based on the
guideline PRISMA statement. The aim of the literature review was
to verify if there were any previous studies that had already
examined the association between an alteration of the nutritional
state and the severity of the disease. The analysed articles [1,18–23]
identify malnutrition as a possible risk factor for COVID-19, but, due
to the lack of clinical trials and the lack of homogenous large
populations to be sampled, this association needs further research
[22,23].

From an additional literature review, the MUST index was
identified as a primary assessment tool to evaluate the risk of
malnutrition in patients that resulted positive for COVID-19 [24].
This tool is quick and easy to apply, with a sensibility of 79% and a
specificity of 91% [25]. A systematic review [26] that evaluated the
use of different assessment scales for malnutritional risk in COVID-
19 positive patients, came to the conclusion that the MUST index
and Mini Nutritional Assessment have an increased predictive
validity in patients infected with SARS-CoV-2.

The MUST index analyses three criteria: BMI, weight changes in
the past 3–6 month and no nutritional intake or likelihood of no
intake for more than 5 days associated with an acute illness.

2.2. Statistical analysis

Data were described as number and percentage or mean and
standard deviation or median and interquartile range, as appro-
priated. Adherence to Gaussian distribution was verified with
Shapiro–Wilks test. Differences among groups were explored with
Kruskal Wallis test or ANOVA, as appropriate. Due to the fact that
MUST index was not calculated on obese patients, a subgroup
analysis was performed in obese and non-obese patients.

Association with mortality was explored with Cox regression.
Independent variables with a p-value under 0.2 were then sub-
mited to a multivariable Cox regression. A p-value under 0.05 was
considered significant. All analyses were performed with Stata 15.

Demographic data and outcome measures were collected in an
excel sheet in anonymous form and collected by an investigator not
involved in assistance, to promote independence in data collection.

The dataset was subdivided in four sections: admission data
(gender, age, BMI, vital sign collected in ER, fasting, enteral
nutrition activation, weight variation, previous pathologies),
MUST value retrospectively calculated by the investigator, hospi-
talisation data (decubitus lesions, superinfections, oxygen ther-
apy, ICU admission), and discharge data (hospitalisation length,
patient status at discharge).

As for superinfections patients diagnosed with positive blood
cultures, pharyngeal swabs and bronchial aspirate were taken into
consideration.

The MUST index score was calculated based on the data given
on the patient record [25]. Factors that affect the patient for fasting
such as: C-PAP, non-invasive ventilation, oxygen masks with an
oxygen flow over 10 L/min and Venturi mask with FiO2 over 50%
were taken into consideration.

Patients were initially divided into 3 different cohorts: patients
that had all hospitalization data documented (n = 529), patients
without a measured BMI (n = 96) and patients for whom it was not
possible to document the weight variations (n = 27). Between
groups comparison has detected statistically significant differences
for age (respectively 65.1 ± 14.3 years, vs 73.3 ± 14.2 years; vs
68.3 ± 12.7 years; p < 0.001), enteral nutrition (respectively 18
(3.4%), vs 12 (12.5%), vs 4 (14.8%), p < 0.001) and intubation at
admission (respectively 11 (2.1%), vs 11 (11.5%), vs 4 (14.8%),
p < 0.001).

The analysis was limited to 515 patients, from whom 353 (68.5%)
males, with also discharge complete information. Examining this
cohort, we estimated the prevalence of malnutrition using the
MUST index, showing an average risk of malnutrition (MUST
½) in 80 (15.5%) patients, 13 (2.5%) patients and higher risk (MUST≥2)
in 80 (15.5%) patients, for a total of patients at risk of 18.1%. The data were analysed
dividing the cohort based on the outcome of the disease (Table 1). The multivariable analysis shows that the association between the
MUST index with mortality is maintained (HR 1.25, 95% CI
1.04–1.51, p = 0.019) even after the corrections of age, treated hy-
pertension, admission to an intensive care unit and oxygen therapy.

Using the same cohort, we analysed the data for morbidity at
discharge: from 515 patients with clinical outcome data at
discharge, 358 were discharged, 65 were transferred to other care
facilities due to clinical problems and 92 deceased (Table 2). The data show an association with the MUST index score (discharged
0.27 ± 0.68, discharged with problems 0.40 ± 0.93, deceased
Table 1
Association of survival with baseline characteristics of the patients.

|                        | Alive | Deceased | HR (95%CI) | p     |
|------------------------|-------|----------|------------|-------|
| N                      | 423   | 92       |            |       |
| Sex (M)                | 288   | 65       | 0.97 (0.62–1.52) | 0.899 |
| Age (years)            | 62.5±13.8 | 76.9±9.7 | 1.07 (1.05–1.09) | <0.001 |
| BMI                    | 27.4±5.2 | 27.0±6.7 | 0.98 (0.94–1.02) | 0.278 |
| Fasting                | 44 (10.4%) | 28 (30.4%) | 2.71 (1.73–4.23) | <0.001 |
| Oxygen Therapy         | 214 (50.6%) | 69 (75.0%) | 2.13 (1.33–3.43) | 0.002 |
| Intubation at T0       | 3 (0.7%) | 7 (7.6%) | 3.55 (1.64–7.69) | 0.001 |
| Dyspnoea               | 216 (51.1%) | 63 (68.5%) | 1.88 (1.21–2.92) | 0.005 |
| Arterial hypertension in therapy | 179 (42.3%) | 69 (75.0%) | 2.99 (1.86–4.79) | <0.001 |
| Current neoplasia      | 22 (5.2%) | 19 (20.7%) | 3.28 (1.98–5.45) | <0.001 |
| MUST                   | 0.29±0.73 | 0.64±0.93 | 1.38 (1.14–1.67) | 0.001 |
| ICU                    | 69 (16.3%) | 10 (10.9%) | 0.31 (0.16–0.61) | 0.001 |

Table 2
Comparison of discharged patients, discharged patients with problems and deceased patients.

|                        | Discharged | Discharged with problems | Deceased | p     |
|------------------------|------------|--------------------------|----------|-------|
| N                      | 358        | 65                       | 92       |       |
| Sex (M)                | 238 (66.5%) | 50 (76.9%) | 65 (70.7%) | 0.222 |
| Age (years)            | 61.1±13.5 | 70.0±13.4 | 76.9±9.7 | <0.001 |
| BMI                    | 27.5±5.1 | 27.1±5.4 | 27.0±6.7 | 0.160 |
| Fasting                | 34 (9.5%) | 10 (15.4%) | 28 (30.4%) | <0.001 |
| Ongoing enteral nutrition | 6 (1.7%) | 2 (3.1%) | 8 (8.7%) | 0.004 |
| Intubation at T0       | 3 (0.8%) | 0 | 7 (7.6%) | 0.001 |
| Dysphagia              | 4 (1.1%) | 7 (10.8%) | 4 (4.8%) | <0.001 |
| Arterial hypertension in therapy | 141 (39.4%) | 38 (58.5%) | 69 (75.0%) | <0.001 |
| Current neoplasia      | 18 (5.0%) | 4 (6.2%) | 19 (20.7%) | <0.001 |
| MUST                   | 0.27±0.68 | 0.40±0.93 | 0.64±0.93 | <0.001 |
| Pressure sores         | 15 (4.2%) | 8 (12.3%) | 3 (3.3%) | 0.012 |
| Superinfections        | 70 (19.6%) | 19 (29.2%) | 20 (21.7%) | 0.211 |
| Oxygen Therapy         | 237 (66.2%) | 54 (83.1%) | 84 (91.3%) | <0.001 |
| NIV-Cpap               | 48 (13.4%) | 18 (27.7%) | 26 (28.3%) | <0.001 |
| Hospitalisation days   | 13 (2–66) | 26 (2–59) | 14 (7–38) | 0.033 |
| Hospitalisation length | 11 (3–91) | 17 (4–80) | 8 (2–69) | <0.001 |

0.64 ± 0.93, p < 0.001). The MUST index was also associated with complications during hospitalisation. In this case, data indicated that the malnutrition index is not associated with superinfection (OR 1.13, 95% CI 0.88–1.47, p = 0.339), nor with development of pressure sores (OR 1.25, 95% CI 0.81–1.64, ±0.64). The development of pressure sores and superinfections appears to be directly related to the length of the hospitalisation and the clinical status of the patient (Table 3).

A subgroup analysis was performed on 387 patients, of which 270 (69.8%) males with BMI<30, divided on the outcome alive/deceased. In this analysis the MUST index score was associated with the survival of the patient (HR 1.43, 95% CI (1.17–1.74), p < 0.001). Lastly, we evaluated the data for morbidity at discharge in this subgroup. These patients also showed a statistically significant association of MUST index score and clinical outcome at discharge (data not showed).

Table 3
Secondary outcome (pressure sores and superinfections).

|                        | Pressure Sores | Superinfections | p     |
|------------------------|----------------|-----------------|-------|
|                        | Present        | Absent          |       |
|                        | 26             | 489             |       |
| Sex (M)                | 19 (73.1%)     | 334 (68.3%)     | 0.672 |
| Age (years)            | 71.9±13.7      | 64.7±14.2       | 0.009 |
| BMI                    | 26.6±4.2       | 27.4±5.6        | 0.735 |
| Fasting                | 5 (19.2%)      | 67 (13.7%)      | 0.390 |
| Ongoing enteral nutrition | 0             | 16 (3.3%)       | 1.000 |
| Oxygen Therapy         | 20 (76.9%)     | 263 (53.8%)     | 0.025 |
| Intubation at T0       | 0              | 10 (2.0%)       | 1.000 |
| Dysphagia              | 2 (7.7%)       | 13 (2.7%)       | 0.172 |
| Arterial hypertension in therapy | 17 (65.4%) | 231 (47.2%) | 0.106 |
| Current neoplasia      | 0              | 41 (8.4%)       | 0.253 |
| MUST                   | 0.34±0.77      | 0.50±0.86       | 0.243 |
| Oxygen Therapy         | 24 (92.3%)     | 351 (71.8%)     | 0.022 |
| NIV-Cpap               | 8 (30.8%)      | 84 (17.2%)      | 0.109 |
| Hospitalisation days   | 19.5 (16–47)   | 13 (2–66)       | 0.022 |
| Hospitalisation length | 36 (6–74)      | 11 (2–91)       | <0.001 |
**In the subgroup of the 128 obese patients, from whom 83 (64.8%) male, the most relevant indicator for outcome from a clinical point of view is the BMI, being directly proportional to mortality.**

In patients with a BMI > 30 transferred to other care facilities mortality was associated with age, BMI, intubation, oxygen therapy, dysphagia, as well as an increase in the average length of hospitalisation.

4. Discussion

This study was carried out with the aim of investigating the association between the risk of malnutrition recorded in the Emergency Department and patients hospitalised due to COVID-19, the clinical outcomes of hospitalisation and the morbidity at discharge.

As for the evaluation of the nutritional risk, identified with the MUST index, the data show a close association between the risk score and the gravity of the disease. Therefore, the MUST index could forecast the outcome of the hospitalisation, regardless of the factors named above.

The average score of the MUST index has shown to be higher in patients that were discharged with clinical problems or that deceased, in line with findings in literature, but our sample has a mean age of 65.1 ± 14.3 years, unlike other studies previously published in the literature [1,2,26] which included a sample with higher ages and lower sample numbers, also our sample presented a lower malnutrition risk (an average value in all groups below 1).

Just as in previous studies [27] our study highlighted that arterial hypertension, suffering from a neoplastic disease or hospitalisation in an intensive care unit are risk factors for severe clinical outcomes. The secondary analysis of obese patients showed an increase in mortality risk directly linked to an increase in age and BMI, underlying diseases like arterial hypertension, diabetes or renal failure were also taken into consideration. These outcomes are confirmed by various articles found in literature that examined this association [28]. This demonstrates that obese patients are at higher risk of a severe evolution of SARS-CoV2 infection and its associated mortality. A limit of this study is the retrospective nature of the analysis. Another limitation to the study is the fact that the analysis was only carried out on a specific subgroup not representative of the general population, as much the analysed groups patient data available differed. Therefore, the results can only be generalised to a limited extent.

Further studies, possibly prospective researches, are therefore needed to obtain a better comprehension of the impact of malnutrition on the outcome of hospitalised COVID-19 patients.

5. Conclusions

Based on the indications given by literature, this study has shown how malnutritional risk is a negative prognostic factor in terms of mortality, hospitalisation length and clinical status of the patient at discharge in patients that have COVID-19.

### Availability of the data and materials

The analysed data is available on demand.

### Ethical approval and consent on the participation

Protocol number: CLI 20/02 from 20/05/2020.

### Sources of funding

The authors declare that there are no sources of funding, neither from internal bodies nor external bodies.

### Authors contributions

Stefano Mancin: Conceptualization, Methodology, Writing — Review & Editing, Investigation, Visualization.

Daniela Cattani: Conceptualization, Methodology, Writing — Review & Editing, Investigation, Visualization.

Andrea Bertone: Writing — Original Draft, Investigation, Resources, Visualization.

Emanuela Morenghi: Software, Validation, Formal analysis, Data Curation, Visualization.

Beatrice Mazzoleni: Review & Editing, Project administration, Supervision.

Elena Azzolini: Review & Editing, Supervision.

Daniela Donizzetti: Supervision.

Lorena Passadori: Project administration, Supervision.

Fanny Sökeland: Writing — Review & Editing. All authors read and approved the final manuscript.

### Declaration of competing interest

The authors declare the absence of any kind of conflict of interest.

### References

[1] Li T, Zhang Y, Gong C, Wang J, Liu B, Shi L, et al. Prevalence of malnutrition and analysis of related factors in elderly patients with COVID-19 in Wuhan, China. Eur J Clin Nutr 2020. https://doi.org/10.1038/s41430-020-0642-3. Apr 22.

[2] Rivellese A, Annucci G, Capaldo B, Vaccaro O, Riccardi G. Nutrizione Umana. Idelson-Gnocchi. 2017:42–4.

[3] Ansero M, Ventriglia G. La malnutrizione “per difetto”: un problema sottostimato? Medea; 2011. p. 55–62.

[4] Alwarawrah Y, Kienann K, MacVeer NJ. Changes in nutritional status impact immune cell metabolism and function. Front Immunol 2018. https://doi.org/10.3389/fimmu.2018.01655.

[5] Najera O, Gonzalez C, Toledo G, Lopez L, Rocio O. Flow cytometry study of lymphocyte subsets in malnourished and well-nourished children with bacterial infections. Clin Diag Lab Immunol 2004. https://doi.org/10.1128/CDLI.11.3.577-580.2004.

[6] Procaccini C, De Rosa V, Galgani M, Carbone F, Cassano S, Greco D, et al. Leptin-induced mTOR activation defines a specific molecular and transcriptional signature controlling CD4+ effector T cell responses. Immuno 2012;189:2941–53. https://doi.org/10.1002/jimm.2200935.

[7] Hoshino Y, Psenicak L, Cohen J, Strauss E S. Rates of reactivation of latent herpes simplex virus from mouse trigeminal ganglia ex vivo correlate directly with viral load and inversely with number of infiltrating CD8+ T cells. J Virol 2007;81:8157–64. https://doi.org/10.1128/JVI.00474-07.

[8] Mora JR, Iwata M, von Andrian UH. Vitamin effects on the immune system: vitamins A and D take centre stage. Nat Rev Immunol 2008;8:685–98. https://doi.org/10.1038/nri2578.

[9] Shirakawa AK, Nagahubo D, Hieshma K, Nakayama T, Jin Z, Yoshih O. 1,25-dihydroxyvitamin D3 induces CCR10 expression in terminally differentiating human B cells. J Immunol 2008;180:2786–95. https://doi.org/10.4049/jimmunol.180.5.2786.

[10] Gombar AF, Pierre A, Maggini S. A review of micronutrients and the immune system-working in harmony to reduce the risk of infection. Nutrients 2020;12:236. https://doi.org/10.3390/nu1210236.

[11] Briguglio M, Pregliasco FE, Lombardi G, Perazzo P, Ban G. The malnutritional status of the host as a virulence factor for new coronavirus SARS-CoV-2. Front Med 2020;7:146. https://doi.org/10.3389/fmed.2020.00146.

[12] Hibbert K, Rice M, Malhotra A. Obesity and ARDS. Chest 2012 Sep;142(3): 785–90. https://doi.org/10.1378/chest.12-0117.

[13] Louie JK, Acosta M, Samuel MC, Schechter R, Vugia D J, Harriman K. A novel risk factor for a novel virus: obesity and 2009 pandemic influenza A (H1N1). Clin Infect Dis 2011;52:301–12. https://doi.org/10.4049/cid.11q52.

[14] Mikiko W, Risi R, Tuccinardi D. Obesity and SARS-CoV-2: a population to safeguard. Diabetes Metab Res Rev 2020;36:e3325. https://doi.org/10.1002/dmrr.3325.

[15] Sharma JR, Yadav UCS. COVID-19 severity in obese patients: potential mechanisms and molecular targets for clinical intervention. Obes Clin Nutr 2021. https://doi.org/10.1093/clin/nca073.
[16] Patel VB, Basu R, Oudit GY. ACE2/Ang 1-7 axis: a critical regulator of epicardial adipose tissue inflammation and cardiac dysfunction in obesity. Adipocyte 2016;5(3):306–11.

[17] Pironi L, Sasdelli A, Ravaoli F, Baracco B, Battaiola C, Bocedì G, et al. Malnutrition and nutritional therapy in patients with SARS-CoV-2 disease. Clin Nutr 2020;40(3):1330–7. https://doi.org/10.1016/j.clnu.2020.08.021.

[18] Briguglio M, Pregliasco FE, Lombardi G, Perazzo P, Banfi G. The malnutritional status of the host as a virulence factor for new coronavirus SARS-CoV-2. Front Med 2020;7:146. https://doi.org/10.3389/fmed.2020.00146.

[19] Boudad L, Lescure F, Lucet J, Yazdanpanah Y, Timsit J F. Severe SARS-CoV-2 infections: practical considerations and management strategy for intensivists. Intensive Care Med 2020. https://doi.org/10.1007/s00134-020-05967-x. Feb 26.

[20] Li Tao, Zhang Y, Gong C, Wang J, Liu B, Shi L, et al. Prevalence of malnutrition and analysis of related factors in elderly patients with COVID-19 in Wuhan, China. Eur J Clin Nutr 2020;74:871–5. https://doi.org/10.1038/s41430-020-0942-3.

[21] Heloneida A, Aquino J, da Silva-Maya J, de Lima Vale H S, Leal Lima Maciel B, Sousa Passos T. Nutritional status, diet and viral respiratory infections: perspectives for severe acute respiratory syndrome coronavirus. Br J Nutr 2020. https://doi.org/10.1017/S0007114520001311.

[22] James PT, Ali Z, Armitage A, Bonell A, Cerami C, Drakesmith H, et al. The Role of Nutrition in COVID-19 Susceptibility and Severity of Disease: A Systematic Review. J Nutr 2021;151(7):1854–78. https://doi.org/10.1093/jn/nxab059.

[23] Popkin B, Du S, Green D W, Beck A M, Algaith T, Herbst H C, et al. Individuals with obesity and COVID-19: a global perspective on the epidemiology and biological relationships. Obes Rev 2020;21(11 November 2020). https://doi.org/10.1111/obr.13126.

[24] Vivek SM, Khaiwal R, Savita V, Bhadada K S, Singh M. Higher body mass index is an important risk factor in COVID-19 patients: a systematic review and meta-analysis. Environ Sci Pollut Control Ser 2020. https://doi.org/10.1007/s11356-020-10132-4.

[25] Practical guidance for using ‘MUST’ to identify malnutrition during the COVID-19 pandemic Malnutrition Action Group (MAG). BAPEN 2020. at, https://www.bapen.org.uk/pdfs/covid-19/covid-mag-update-may-2020.pdf. [Accessed 4 April 2020].

[26] Marinos E. The “MUST report”. Nutritional screening of adults: a multidisciplinary responsibility. BAPEN 2020.

[27] Oliveira S, David F. Nutritional risk screening tools for older adults with COVID-19: a systematic review. Nutrients 2020;12:2956. https://doi.org/10.3390/nu12102956.

[28] Yang J, Zheng Y, Gou X. Prevalence of comorbidities and its effects in patients infected with SARS-CoV-2: a systematic review and meta-analysis. Int J Infect Dis 2020;94:91–5.