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Protocol to Determine the Extent of Lung
Injury in COVID-19 Pneumonia
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Giovanni Volpicelli, MD FCCP; Thomas Fraccalini, MD; Luciano Cardinale, MD; Giuseppe Stranieri, MD;
Rouslan Senkeev, MD; Guido Maggiani, MD; Alberto Pacielli, MD; and Domenico Basile, MD
BACKGROUND: Lung ultrasound (LUS) scanning is useful to diagnose and assess the severit
of pulmonary lesions during COVID-19-related ARDS (CoARDS). A conventional LUS scor
is proposed to measure the loss of aeration during CoARDS. However, this score was val
dated during the pre-COVID-19 era in patients with ARDS in the ICU and does not conside
the differences with CoARDS. An alternative LUS method is based on grading the percentag
of extension of the typical signs of COVID-19 pneumonia on the lung surface (LUSext).
RESEARCH QUESTION: Is LUSext feasible in patients with COVID-19 at the onset of disease
and does it correlate with the volumetric measure of severity of COVID-19 pneumonia le
sions at CT scan (CTvol)?
STUDY DESIGN AND METHODS: This observational study enrolled a convenience sampling of
patients in the ED with confirmed COVID-19 whose condition demonstrated pneumonia a
bedside LUS and CT scan. LUSext was visually quantified. All CT scan studies were analyze
retrospectively by a specifically designed software to calculate the CIvol. The correlation
calculated.
RESULTS: We analyzed data from 179 patients. Feasibility of LUSext was 100%. Time t
perform LUS scan was 5 \pm 1.5 mins. LUSext and CTvol were correlated positively ($R=0.6$
P < .0001). Both LUSext and CTvol showed negative correlation with Pao ₂ /Fio ₂ ratio ($R =$
-0.66 and $R = -0.54$; $P < .0001$, respectively).
INTERPRETATION: LUSext is a valid measure of the severity of the lesions when compare
with the CT scan. Not only are LUSext and CTvol correlated, but they also have simila
inverse correlation with the severity of respiratory failure. LUSext is a practical and simple
bedside measure of the severity of pneumonia in CoARDS, whose clinical and prognost
Impact need to be investigated further. CHEST 2022; CHEST 2022 ; CHEST 202 ; CHE
KEY WORDS: ARDS; COVID-19 pneumonia; CT volumetry; interstitial pneumonia; lun
ultrasound scan; lung ultrasound scoring
ABBREVIATIONS: CoARDS = corona virus-related ARDS; CTvol = CT Medical Sciences (G. M.), Section of Geriatrics, Città della Salute e del
volumetric scoring; GGO = ground glass opacities; LUS = lung ultra- sound scorping, LUS et = lung ultracound extension scoring, P/E =

AFFILIATIONS: From the Departments of Emergency Medicine (G. V. and T. F.), and Oncology (L. C., G. S., R. S., A. P., and D. B.), Radiology 54 **Q2** Unit, San Luigi Gonzaga University Hospital; and the Department of

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Take-home Points

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Study question: Is a simplified lung ultrasound scoring, based on the eyeball estimation of the extension of pulmonary lesions (LUSext), accurate to assess the severity of lung injury in COVID-19 pneumonia?

Results: In patients with first diagnosis of COVID-19 pneumonia, LUSext correlated positively with the volumetric measure of lung lesions at CT scan (R = 0.67; P < .0001) and showed a negative correlation with the Pao₂/Fio₂ ratio (R = -0.66; P < .0001).

Interpretation: A simple eyeball estimation of the percentage of extension on the chest surface of the typical sonographic lesions can be used to measure the lung damage during the first diagnostic approach to COVID-19 pneumonia.

131 Chest imaging is strategic not only in the initial 132 diagnosis of COVID-19-associated pneumonia but also 133 to evaluate and monitor its severity. The diagnostic role 134 of lung ultrasound scanning (LUS) has been explored 135 136 widely and compared with confirmation by CT scan and 137 reverse transcriptase-polymerase chain reaction (RT-138 PCR).¹⁻³ LUS diagnosis of COVID-19 pneumonia is 139 based on the recognition of typical interstitial and 140 parenchymal signs, together with the careful evaluation 141 of their distribution on the lung surface and 142 combination with clinical phenotypes.^{1,2} 143

146 Study Design and Methods

147 Temporal Framework

We performed an observational cross-sectional single-center study on
a convenience sampling of patients with confirmed first diagnosis of
COVID-19 pneumonia. Patients were enrolled in the ED of San
Luigi Gonzaga University Hospital during the first wave (March to
May 2020) and the second wave (October 2020 to February 2021) of
the COVID-19 pandemic.

¹⁵⁴ Selection of the Population

155 During the study period in our ED, all patients suspected of COVID-19 156 were examined by LUS at their first visit. Consecutive patients for 157 whom CT scans were performed immediately after LUS for clinical reasons independent from the study protocol, were considered 158 eligible. Only adult patients with final demonstration of acute 159 COVID-19 pneumonia with both positive chest imaging and positive 160 RT-PCR were selected. Based on a convenience sampling, those 161 patients who were examined for LUSext scoring by specifically 162 trained operators entered the analysis. Information about symptoms of presentation, the timing of symptoms onset, and first bedside 163 clinical data, which included the Pao2/Fio2 ratio (P/F), was recorded 164 systematically. Patients were grouped in three different clinical 165 phenotypes at presentation, according to a protocol previously

166 Together with the first diagnosis, the LUS quantification 167 of the severity of COVID-19 pneumonia might be of 168 great use in the management of the disease and its 169 prognostication. The advantage of the use of LUS is 170 based on its high feasibility and the possibility to repeat 171 the examination at the bedside without the necessity to 172 move the patient, thus reducing the possibility of 173 intrahospital cross infection during a pandemic surge. 174

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LUS is a surface imaging technique, quite limited in the evaluation of lesions that do not abut the lung periphery.^{4,5} However, COVID-19 typically affects mainly the lung periphery and is characterized by typical lesions alternating with spared areas. Thus, a possibility to assess the severity of COVID-19 pneumonia is to assign a LUS score based on the visual estimation of the percentage of extension of the typical lesions on the pulmonary surface (LUSext).

Our hypothesis is that the intrinsic characteristics of COVID-19 pneumonia makes LUSext a reliable tool in the assessment of the severity of the lung damage during the first approach to the disease. The primary aim of our study is to investigate the correlation between the LUSext score and the volumetric assessment of pulmonary lesions calculated by CT scan (CTvol) in patients with COVID-19 pneumonia. A secondary aim was to evaluate whether the severity of COVID-19 pneumonia assessed by LUSext and CTvol correlates with the objective measure of respiratory failure.

validated and based on the presence of preexisting chronic cardiac or respiratory diseases (mixed phenotype) and on the presence (severe phenotype) or absence (mild phenotype) of signs and/or symptoms of respiratory failure.^{1,3} The list of significant chronic conditions included severe COPD, pulmonary fibrosis, lung cancer, heart failure, or cor pulmonale. Respiratory failure was determined according to the presence of dyspnea, either objective or self-reported, and/or desaturation after walking and/or demonstration of P/F < 300 mm Hg. The local Ethical Committee approved the Q5 protocol of this study (Registro di Protocollo Generale n°2840 - 210221).

Lung Ultrasound Scan

A complete LUS examination was performed at presentation on the 212 anterior, lateral, and posterior chest, as previously described.^{1,2,6} 213 LUSext was performed by operators specifically trained in the study 214 protocol. Commercially available ultrasound equipment (Mindray 215 TE7; Esaote MyLabSeven) with convex transducers (3.5 to 6.0 MHz) 216 were used. The focus was placed at the height of the pleural line. Depth was set at approximately 8 to 10 cm, according to patient's 217 size. Gain was regulated to optimize the whole image. The 218 sonographers were ED clinicians, with documented experience in 219 using LUS in emergency and critical care. During the examination, 220 the LUS operator was blind to the result of CT scan and RT-PCR

test, but not to the patients' clinical condition. LUS was performed
immediately at presentation whenever possible and always before the
result of the CT scan and the RT-PCR test. Time for LUS
examination was recorded and reported on the data sheet by the
same operator.

LUS diagnosis: Each LUS examination was classified according to standardized, mutually exclusive patterns, already described and validated in an international multicenter study.³ Only patients whose condition showed the following two positive patterns were considered for the study analysis:

High Probability Pattern: Typical LUS pattern of COVID-19
pneumonia has bilateral and multifocal clusters of separated or
coalescent B-lines, large hyperechoic bands (light beams), multifocal
peripheral consolidations, regular and irregular pleural lines, with or
without large consolidations. These clusters should appear in a
patchy distribution, abruptly alternating with normal A-lines
patterns ("spared areas").

Intermediate Probability Pattern: Less typical pattern includes
 unilateral isolated clusters of B-lines and light beam or focal multiple
 B-lines, with or without small peripheral consolidations.

LUS extension score (LUSext): A measure of the superficial extension 240 on the chest wall of the typical COVID-19 LUS signs was calculated 241 during the examination. The visual estimate of the percentage of 242 extension of the lesions was reported by the operator directly at bedside. The examination was performed by recording a video of the following four areas per side (Fig 1): (1) anterior chest in longitudinal scan, (2) lateral chest in longitudinal scan, (3) posterior chest paravertebral in longitudinal scan, and (4) posterior chest below the scapula in oblique scan. In each area, it is possible to examine a variable number of intercostal spaces; usually, there are approximately four spaces in areas 1, 2 and 4, and six or more spaces in area 3. We assigned the score based on the visual extension of lesions in fixed percentages, assigning 0%, 25%, 50%, 75%, or 100% depending on the number of intercostal spaces that show the pathologic signs (Fig 2). We considered the following typical signs of COVID-19 pneumonia to assign the LUS scoring: multiple B-lines separated and coalescent, the "light beam," small peripheral consolidations with irregularity of the pleural line, and large consolidations.³ Absence of any pathologic sign was assigned as 0%. The percentages assigned to each area were then summed and divided by the total 8 areas. This simple technique allows a rough

estimate of the extension of the typical COVID-19 lesions on the 276 lung surface, without the necessity to differentiate B-lines, 277 consolidations, and pleural line characteristics. A demonstrative case 278 is added as supplementary material (e-Video 1-9 and e-Fig E-1). 279

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CT scan

CT examinations were acquired by 64- or 128-bank CT machine ²⁸¹ (Philips Ingenuity and GE) by volumetric chest scans (slice thickness 282 1.25 mm; slice interval 1 mm) with parenchymal lung retro- 283 reconstruction algorithm. Only patients with clinical indication to 284 CT scan, which was decided independently by the physician in 285 charge, were moved to the radiology unit inside the ED to perform the examination. The patients were in supine and head-first position ²⁸⁶ and received scanning with breath held. The following parameters 287 were used: 10 kV; 100 mAs real-time adaptive control; layer 288 thickness 1 to 2.5 mm; pitch, 1 to 1.5; matrix, 512 \times 512. All images $\frac{96}{289}$ were transmitted to the postprocessing workstation and 290 reconstructed with the use of high-resolution and conventional algorithms. Each study was read and interpreted by two expert 291 radiologists with long-standing experience in chest imaging. 292

CT Scan Diagnosis: Signs and nomenclature of CT scan were those recommended for COVID-19 and reported in European and North American societal recommendations.^{7,8} Specifically, we evaluated 295 only patients with confirmation of COVID-19 pneumonia that was supported by the visualization of one of the two CT scan positive readings: (1) "typical appearance" in the presence of peripheral, bilateral, multifocal ground glass opacities (GGO) with or without consolidation or visible lines ("crazy-paving") and (2) "indeterminate appearance" in the presence of multifocal perihilar or unilateral GGO with or without consolidation or very small GGO nonrounded or nonperipheral. 302

CT Scan Volumetric Score: All acquired images were processed 303 retrospectively through a semiautomated external software (Thoracic 304 Vcar; General Electric) to quantify the percentage of aerated, GGO, and consolidated lung parenchyma.⁹⁻¹¹ To this end, densitometric thresholds were identified to differentiate these entities: (1) 700 Hounsfield units to differentiate GGO from consolidated lung. These 308 thresholds were identified based on a method validated in literature and readjusted according to the judgment of the experienced radiologist.⁹ CT scans were segmented semiautomatically by the software and then refined by the experienced radiologist as needed. 312

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318 Figure 1 – A and B, The four chest areas that were examined to assess the extension 319 of the pulmonary lesions in patients with 320 COVID-19 pneumonia. A, Patient is first 321 placed in the supine position: area 1 is 322 scanned longitudinally between the sternum and the anterior axillary line; area 2 323 is scanned longitudinally between the 324 anterior axillary line and the posterior 325 axillary line. B, Patient is then turned in the lateral decubitus: area 3 is scanned 326 longitudinally between the spine and the 327 medial margin of the scapula; area 4 is 328 scanned in oblique (along the intercostal spaces) below the inferior margin of the 329 scapula. The same procedure is then 330 repeated on the other side.

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333 visible by lung ul	trasound examination on							
334 the chest wall. Ea	ach area is examined, and 0-25-50-75-100% is		EXTENSION OF THE CHEST SURFACE (%)					
assigned visually.	The final score in per-		and	0	Antes			
337 centage is given a	by the sum of the per- rea divided for the total of							
338 eight scans.			4	25				
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The software automatically eliminates the airways down to the segmental branches. The vascular volume, estimated to be approximately 3%, was subtracted manually from the percentage of consolidated parenchyma. CT scans with significant motion artifacts and all those performed in patients with significant prior pulmonary alterations (severe interstitial disease, marked emphysema, lung cancer, fibrothorax) and major thoracic deformities were excluded.

381 RT-PCR Swab Test

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The diagnosis of COVID-19 was confirmed by an RT-PCR nasal-pharyngeal and/or bronchial swab (BD SARS-CoV-2 Reagents for
BD MAX System). In pretriage, a hand-reading Rapid Antigenic Test
(COVID-19 Ag Rapid Test Device; Abbott Panbio) or a facilitated

reading (LumiraDx SARS-CoV-2 Ag test; LumiraDx) was performed to guide the first allocation of the patients in different areas of the ED. However, the infection was confirmed only after the RT-PCR swab detailed earlier.

Statistical Analysis

Data are expressed as mean \pm SD. Normality of data was checked with the
use of the Q-Q plot evaluation and Shapiro-Wilk test. Spearman rank
correlation was used to examine the significance of correlation between
variables. Durbin-Watson Test was used to rule out autocorrelation of
residuals, and Breusch-Pagan Test was used to evaluate
heteroscedasticity of residuals variance. Linear regressions were
performed where appropriate. All analyses were calculated with R Studio.435
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441 Results

442 443 Patients

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We analyzed by convenience sampling LUSext and
CTvol of 179 patients with confirmed COVID-19
pneumonia. Sampling was determined by the availability
of the personnel trained in LUSext. Figure 3 shows the
patients' flow of the study; Table 1 shows the patients'
characteristics.

LUS and CT Scan Diagnoses

In all patients, LUS was performed to assign the 453 probability of COVID-19 pneumonia and to calculate 454 455 LUSext. The feasibility of LUS was 100%. Average time 456 spent for the entire LUS examination and the calculation 457 of the LUSext score was 5 ± 1.5 min. In 12 patients (7%) 458 the LUS diagnosis for COVID-19 pneumonia was 459 intermediate probability and high probability in 167 460 patients (93%). CT scan was indeterminate in six 461 patients (3%) and typical in 173 patients (97%). In six 462 patients LUS intermediate probability corresponded to 463 typical CT scan. The other six LUS diagnoses of 464 intermediate probability corresponded to indeterminate 465 at CT scan. All the 167 LUS high probability were typical 466 467 at CT scan. 468

469 LUSext, CTvol and P/F

470 LUSext ranged from 3.125% to 78.125%, and CTvol 471 ranged from 6.219% to 91.533%. A statistically 472 significant correlation was found between LUSext and 473 CTvol (*R* [Spearman Rho] = 0.67; *P* <.0001)(Fig 4A), 474 which pointed to a good accordance between the 475 extension of the lung lesions on the chest surface and the 476 volume of the lung injuries in COVID-19 pneumonia. 477 Regarding the adequacy of all the necessary statistical 478 479 criteria, we calculated the linear regression between

496 LUSext and CTvol (R-Square = 0.52), which suggests 497 that the extensions visualized by LUS can be linked 498 adequately through a linear relationship to the volumes 499 obtained by CT scans. A statistically significant inverse 500 correlation was found between LUSext and P/F values 501 (R = -0.66; P < .0001) (Fig 4B), and between CTvol and 502 P/F values (R = -0.54; P < .0001) (Fig 4C). We analyzed $\frac{1}{503}$ data from a subgroup of patients with more severe 504 respiratory failure at presentation, which was selected 505 for P/F value below 300 mm Hg (Table 1). In this 506 subgroup the correlation between LUSext and CTvol 507 was still significant (R = 0.58; P < .0001; and R-Square 508 509 0.41) (Fig 4D).

Discussion

Our data show that, in patients with acute onset of 513 COVID-19 pneumonia, the LUS score based on the 514 estimate of the LUSext is correlated positively with the 515 516 CTvol. Moreover, both LUSext score and CTvol are 517 correlated inversely to the main clinical index of respiratory function, the P/F values. In the subgroup of 518 519 patients with more severe pneumonia who presented 520 with objective respiratory failure characterized by low P/ 521 F values, LUSext and CTvol showed a similar good 522 positive correlation, even in a condition of more 523 complicated, extended, and mixed lung lesions. Thus, 524 LUS performed as well as CT scan in terms of 525 determining severity of COVID-19 lung lesions. 526

LUS is limited strongly in the visualization of the lung527lesions because ultrasound scanning can explore only the529periphery of the lung. Indeed, the alveolar air represents a530barrier to the visualization of the lung parenchyma.531However, it is well-acknowledged that, when the air532content is impaired and the density of the lung is533increased in the periphery, LUS visualizes with high534



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Characteristic	All Patients (N = 179)	Patients with Pao_2/Fio_2 Ratio <300 mm Hg (N = 114)		
Age, y	66 ± 14	69 ± 13		
Male/female, No.	111/68	74/40 7.8 ± 3.7		
Onset of symptoms, d	7.7 ± 4			
Clinical phenotype, No. (%)				
Severe	126 (70)	108 (95)		
Mild	47 (26)	0		
Mixed	6 (3)	6 (5)		
Lung ultrasound scan pattern, No. (%)				
Intermediate	12 (7)	2 (1.8)		
High	167 (93)	112 (98.2)		
CT scan appearance, No. (%)				
Indeterminate	6 (3)	0		
Typical	173 (97)	114 (100)		
Pao ₂ /Fio ₂ ratio, mm Hg	$\textbf{277} \pm \textbf{83}$	228 ± 54		
Lung ultrasound extension score, %	$\textbf{36.4} \pm \textbf{17.8}$	43.5 ± 16.5		
CT volumetry, %	$\textbf{35.0} \pm \textbf{17.0}$	41.0 ± 16.8		
Time for lung ultrasound scan, min	5.1 ± 1.5	5.2 ± 1.5		

TABLE 1 Detient Characteristics 551

574 sensitivity the typical interstitial and consolidative 575 patterns.¹² In most pulmonary conditions, particularly in 576 emergency situations, these LUS ultrasound patterns in 577 combination with the available clinical information, allow 578 for accurate finalization of the diagnostic process with 579 high specificity. For instance, the high sensitivity and 580 specificity of LUS are well-suited for the early diagnostic 581 workup of acute undifferentiated respiratory failure.¹³ 582

583 During the COVID-19 pandemic, LUS demonstrated a 584 high diagnostic accuracy in the diagnosis of interstitial 585 pneumonia related to the Sars-Cov-2 infection.^{3,14} LUS 586 can visualize the early alterations of the disease, 587 588 including the GGO and the consolidations observed by CT scan in the lung periphery of patients with COVID-589 19 pneumonia.¹⁵ Many authors also hypothesized a role 590 591 in the quantification of lung damage during the acute 592 phase of COVID-19.¹⁶⁻¹⁹ The method of LUS scoring 593 that has been advocated for grading the severity of lung 594 involvement in COVID-19 pneumonia was investigated 595 and validated in the pre-COVID era on patients with 596 classic ARDS who received invasive ventilation.²⁰⁻²² This 597 conventional sonographic score is based on the 598 assignment of three grades of incremental loss of 599 aeration through the recognition of signs of progression 600 from separated and coalescent B-lines to consolidation, 601 602 on 12 anterior, lateral, and posterior chest areas.

603 COVID-19 pneumonia complicated by severe 604 respiratory failure can be included fully in the modern 605

syndromic definition of "corona virus-related ARDS" (CoARDS). However, CoARDS presents peculiar characteristics. Indeed, evolution of the disease, histopathologic condition, extension of the lung damage, and even strategies of ventilatory support are different between ARDS and CoARDS.²³ For instance, a typical characteristic of CoARDS is the significant mismatch between the severity of the lung damage and the respiratory condition of the patient, which is not usual in both respiratory and extra-respiratory ARDS.²⁴ Moreover, although in ARDS, there is clear evidence that the LUS scoring is useful to guide treatment, to date, there is no robust demonstration that the severity of the lung damage assessed by LUS in CoARDS may be used in practice to predict the evolution of COVID-19 pneumonia and guide management with different ventilatory strategies.

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The conventional LUS score is based on the 649 differentiation between separated B-lines, coalescent 650 651 B-lines, and consolidations. This task can be technically 652 difficult to perform at bedside in patients with COVID-653 19. Assigning different grades to B-lines and 654 consolidations may become a complicated task in a 655 condition that, by definition, is characterized by clusters 656 of interstitial signs and consolidations abruptly 657 alternating with spared areas in a patchy distribution. 658 Moreover, in a condition of tight mix of various LUS 659 signs, the differentiation between B-lines and 660



Figure 4 – A-D, Correlations between A, lung ultrasound extension score and the CT scan volumetry of pulmonary lesions; B, lung ultrasound extension score and Pao₂/Fio₂ ratio; and C, CT scan volumetry of pulmonary lesions and Pao₂/Fio₂ ratio, in 179 patients with COVID-19 pneumonia. D, Correlations between lung ultrasound extension score and the CT scan volumetry of pulmonary lesions in a subgroup of 114 patients shows Pao₂/Fio₂ ratio <300 mm Hg at presentation. CTvol = CT scan volumetry; LUSext = lung ultrasound extension score; P/F = Pao₂/Fio₂ ratio; vol = volume.

consolidations might lose its importance as an indicator of different degrees of lung aeration.

Thus, in the opinion of these authors, the complexity of the technique and the physiopathologic mismatch between lesions and function are limitations to the practical application of the conventional LUS scoring in COVID-19. Considering these limitations, we theorized a different LUS approach to quantify the lung damage in COVID-19 pneumonia. The proposed approach is based on the estimation of the extension of the lesions and not on grading lung aeration. The extension of pneumonia on the chest surface seems to be a more appropriate index for the evaluation of the characteristics of this new disease. The new LUSext scoring does not need to differentiate different degrees of aeration and is based on a gross visual estimation of extension of lesions, which makes this new technique more immediate and easier to be performed at bedside.

The main limitation of our study is the lack of demonstration of the usefulness and practical implication of LUSext. For instance, we did not collect data on the outcomes and their correlation with LUSext. However, the only aim of the present study was to investigate the feasibility of LUSext and 755 the correlation between LUSext and CTvol in patients 756 with an initial diagnosis of COVID-19 pneumonia. It 757 is highly probable that LUSext represents just a picture of the disease at a given moment that probably, like other chest imaging and respiratory scorings, is not useful in practice to predict evolution 762 and prognosis in severe cases of COVID-19. In our experience, this new LUS scoring might indicate the 764 necessity for a more careful follow-up and hospitalization in patients with a borderline condition 766 and without signs of respiratory failure who show more severe grades of extension of pneumonia. This 768 hypothesis, together with other possible practical applications of LUSext, need a scientific

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771 demonstration by future observational trials and 772 validating studies. 773

Another limitation is the lack of assessment of 774 interoperator variability of LUSext. This may be 775 considered particularly important given the "eveball" 776 characteristic of our method. However, the aim of this 777 778 study was to introduce a new method and assess the 779 correlation with volumetric CT scan. Indeed, the 780 conventional aeration LUS scoring has some subjectivity 781 that was not assessed in the original introductory 782 study.²¹ During a COVID-19 pandemic surge 783 assessment of variability may be particularly challenging 784 for the necessity to limit exposure of the operators. 785

786 During the study period, more than one-half of the 787 eligible patients who received a CT scan were not 788 analyzed (Fig 3). Indeed, the enrollment was on a 789 convenience sampling based on the availability of the 790 trained operator. However, selection was completely 791 random, and patients were enrolled at any time of the 792 day during the pandemic surge in the ED. Moreover, 793 enrolling only patients with indication to CT scan may 794 have influenced selection of more severe grade of lung 795 injury. Potentially, this might limit generalizability of 796 797 our conclusions to minor forms of COVID-19. 798

A minor limitation is the use of P/F calculated in 799 spontaneous breathing as a measure of respiratory 800 failure. Indeed, it is well-acknowledged that the FIO₂ 801

826 percentage extracted during administration of oxygen 827 flux in mask or calculated in room air is less accurate 828 than the true FIO₂ calculated in intubated patients. 829 However, P/F during spontaneous breathing is 830 indicative of the respiratory status during the initial 831 evaluation and represents the important parameter to 832 decide ventilatory treatment. Moreover, most of the 833 patients who presented to the ED during the COVID-19 834 pandemic outbreak did not need intubation. 835

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Finally, in our study a particularly savvy group of clinicians performed the LUS examinations. This may raise concern about the possibility that expertise affects accuracy. However, a previous international multicenter study demonstrated that LUS in COVID-19 may be performed by several operators with different levels of expertise maintaining high accuracy and low variability.³

Interpretation

Our new LUS score that is based on the eyeball estimation of the percentage of extension on the chest surface of the signs of COVID-19 pneumonia correlates positively with the CTvol scan and inversely with the P/ F at the onset of the disease. This new simplified and practical LUS scoring approach is well-suited to grade the pulmonary damage of this new disease at presentation.

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- Author Contributions: G. V. takes the 806 responsibility for the content of the 807 manuscript, including the data and analysis. G. V. designed the study; G. V., T. F., and G. 808 M. contributed equally to perform the 809 ultrasound examinations; L. C., G. S., R. S., A. 810 P., and D. B. performed and interpreted the CT scan studies; G. S., A. P., and D. B. 811 performed the volumetric calculations of the 812 CT scan examinations; R. S. is responsible for 813 the statistical analysis; G. V., T. F., L. C., R. S., and G. M. had full access to all the data in the
- 814 study; G. V., T. F., L. C., and G. M.
- 815 contributed substantially to the writing of the
- 816 manuscript; all authors read and approved the final version of the manuscript.
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