

Pulmonary and Critical Care Medicine

Covid-19 Bootcamp #2

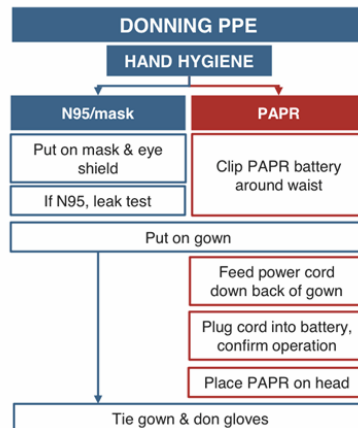
Jasleen Minhas, Katie Auriemma, Roger Kim

March 31, 2020



Presentation Slides and Tip sheets





HIGH-RISK PROCEDURES & ALTERNATIVES	
Nebulized bronchodilators	MDI for COVID+ or PUI
Inhaled epoprostenol (Veletri)	Maximize all other ARDS adjuncts
BIPAP or HFNC	Intubate if impending respiratory failure Do not switch between modalities
Mask ventilation	Rapid seq. intubation
Ventilator disconnect Open suctioning	Minimize if possible

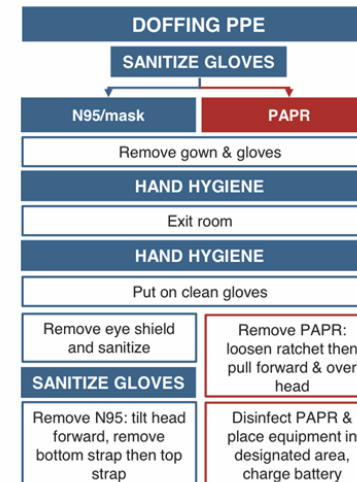
Also includes: intubation, CPR, bronchoscopy

Penn Medicine Tip Sheet: Personal Protective Equipment

Updated 3/25/2020 – [newest version here](#)

PPE GUIDELINES		
Workspaces		Droplet + Social Distancing
Non-PUI		Droplet
PUI	PUI ED	Droplet + Contact
	PUI Ward	Droplet + Contact
	PUI ICU	Airborne* + Contact
Confirmed COVID-19 + *Droplet if PPE/room unavailable		Airborne* + Contact
Aerosol Generating Procedures (PUI or COVID-19+)		Airborne + Contact + Eye Shield

RRT/CODE		
Non-PUI	RRT (respiratory)	Droplet + Contact + Eye Shield
	Code and/or Intubation	Droplet + Contact + Eye Shield
PUI	RRT (any)	Droplet + Contact + Eye Shield
	Code and/or Intubation	Airborne + Contact + Eye Shield
Confirmed COVID-19 +	RRT (any)	Airborne + Contact + Eye Shield
	Code and/or Intubation	Airborne + Contact + Eye Shield



Remove gloves & HAND HYGIENE

CONSERVING PPE

Relocate IV pumps & vent control outside room
Bundle invasive procedures
Consolidate lab draw & med admin times
Order imaging only if results will change care

CHECK SHAREPOINT FOR UPDATES

Penn Medicine Intranet

HOME Service Requests SITE DIRECTORIES

What's Hot COVID-19 UPDATES COVID-19

A quick-reference guide created for the Division of Pulmonary and Critical Care Medicine. Contents may change as situation demands. Email Jeff Min & Jennifer Ginestra for corrections.

- ▶ Please reach out to the following fellows for questions:
- ▶ Jeff Min: jeff.min@pennmedicine.upenn.edu
- ▶ Jen Ginestra: Jennifer.ginestra@pennmedicine.upenn.edu
- ▶ Jasleen Minhas: jasleen.minhas@pennmedicine.upenn.edu

Screening and Diagnosis



HUP COVID-19 Testing - Logistics

- ▶ Rapid Cartridge PCR with run time of < 45 minutes
- ▶ Specimens: obtained by MDs
 - Non intubated: NP or OP swab
 - Intubated: NP swab only
- ▶ Can be added on to RPP if sent in ED
- ▶ In house testing: currently only by approval from ID


Key Contact Information

**HUP ID APPROVAL PAGER: (215)-614-0895 →
contact 24/7 for in-house test approval**

HUP COVID-19 Testing - Logistics

	Daily Volume	Result time	Population targeted	Order
Urgent	120 tests/day	<4 hours	ED/Obs/Inpatients	Golden Ticket
Less Urgent	200 tests/day	4-12 hours	Occ Exposures	Blue Ticket

- As of 3/30 testing is being offered 24 hours a day

 Penn Medicine Hospital of the University of Pennsylvania 3400 Spruce Street, Philadelphia, PA 19104 DIVISION OF LABORATORY MEDICINE Irving Nachamkin, DrPH, MPH Laboratory Director		Patient Label
<input type="checkbox"/> Check box for Occupational Medicine Patient		
Date and Time Collected (required by law) Date: _____ Time: _____	Name of Collector (required by law) Name: _____ (print)	Requesting Provider name Name: _____ (print)
SARS-CoV-2 (COVID-19) Testing Requisition		

HUP COVID19 – Inpatient Protocol

Has patient had a recent RPP or rapid flu/RSV sample that is still in HUP lab

YES

NO

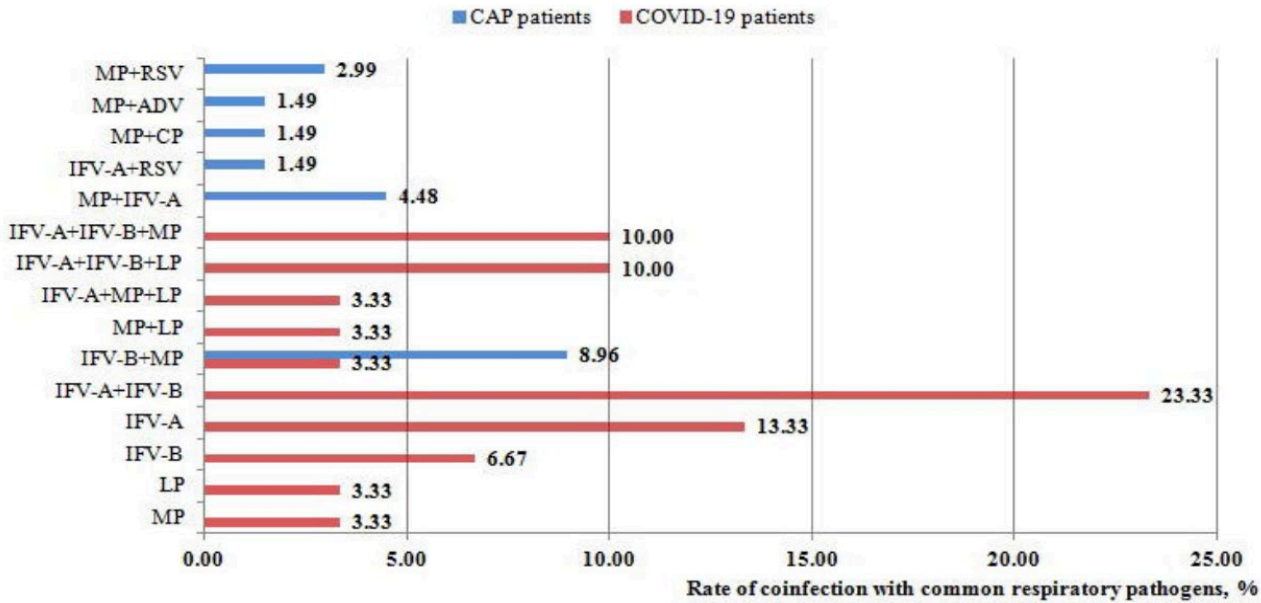
1. Call ID approval pager
2. Infection control completes COVID-19 Requisition form aka “Golden Ticket”.
No order entry in EPIC
3. Deliver / tube form to Central Receiving (Founders 7)

1. Call ID approval pager
2. Infection control completes COVID-19 Requisition form aka “Golden Ticket” – form emailed/delivered in person
3. Collect Specimen
4. Deliver / tube form to Central Receiving (Founders 7)

Co-infections

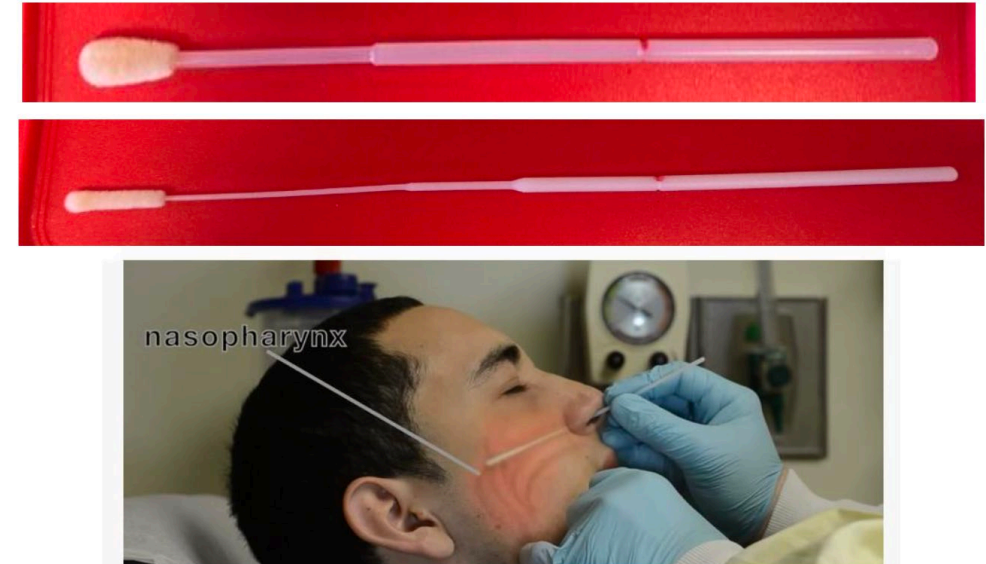
Table 4. Common respiratory pathogens detected in COVID-19 patients and healthy controls

Pathogens detected	COVID-19 patients (n=30)	Healthy controls (n=30)	P-value
	No. (%)	No. (%)	
IFV-A	18 (60.00)	0 (0.00)	<0.0001
IFV-B	16 (53.33)	4 (13.33)	0.0018
MP	7 (23.33)	0 (0.00)	0.0105
LP	6 (20.00)	0 (0.00)	0.0237
RSV	0 (0.00)	0 (0.00)	NA

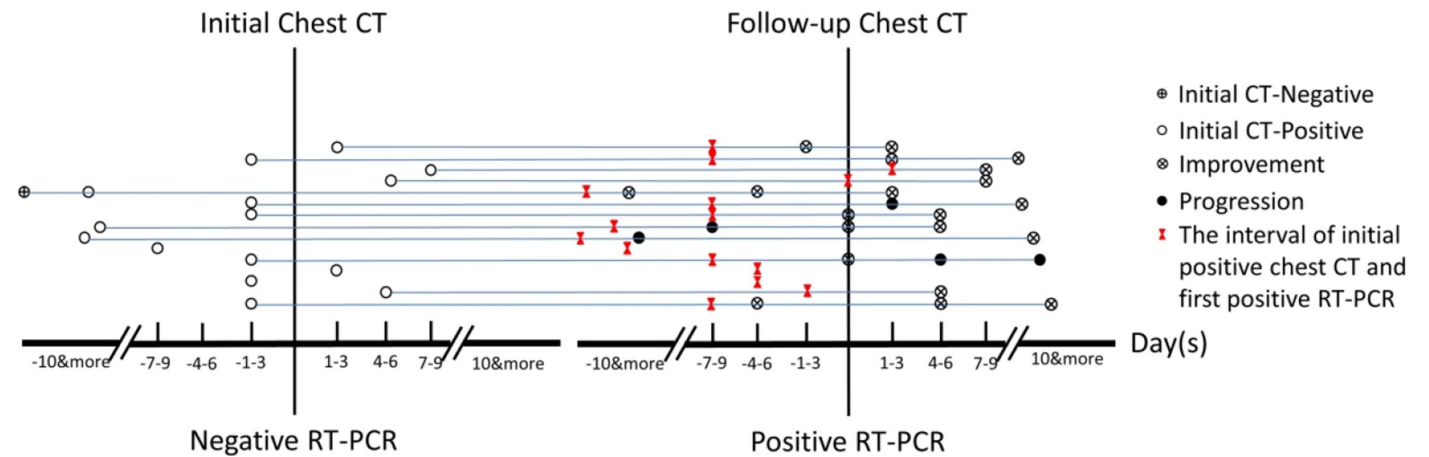
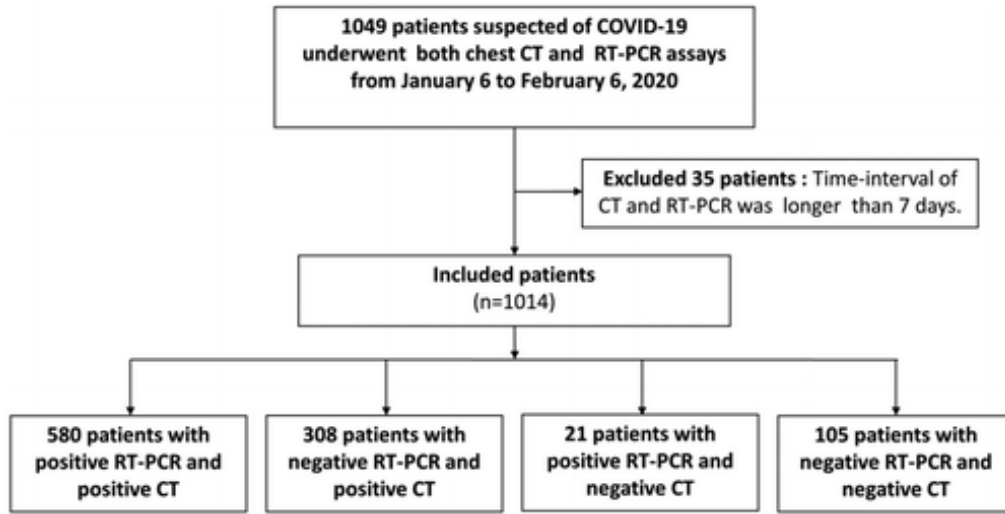


Nasopharyngeal Swabs

- ▶ PPE: surgical mask + eye protection + gown + gloves
- ▶ Procedure:
 - Tilt head back & insert swab parallel to palate
 - Stop when resistance met
 - Leave in place for 2 – 3 seconds
 - Rotate 10 – 15 sec
 - Repeat on other side
- ▶ OP swab: larger swab
- ▶ Immediately place both into sterile viral transport media vial
- ▶ Double bag specimen
- ▶ Include “golden ticket”
- ▶ Deliver to central receiving (Founders 7)



Correlation of Chest CT and RT-PCR



Screening for COVID 19

- ▶ Logistics of screening
- ▶ Co-infections may occur with COVID 19
- ▶ Procedure of obtaining NP swab
- ▶ False negatives – more to come as we learn

Clinical Features and Course

Katie Auriemma



Actual Patients admitted in UPHS System

- ▶ 72M with confusion and a fever
- ▶ 80M with disequilibrium – activated for code stroke
- ▶ 68W with fever and SOB
- ▶ 72M with 1 week of diarrhea and emesis

All Covid-19 Positive



Early Reports of Epidemiology and Clinical Characteristics

Epidemiological and clinical characteristics of 99 cases of 2019 novel coronavirus pneumonia in Wuhan, China: a descriptive study

Nanshan Chen*, Min Zhou*, Xuan Dong*, Jieming Qu*, Fengyun Gong, Yang Han, Yang Qiu, Jingli Wang, Ying Liu, Yuan Wei, Jia'an Xia, Ting Yu, Xinxin Zhang, Li Zhang

- ▶ Lancet, February 15, 2020
- ▶ Largest early cohort of hospitalized patients
- ▶ Fever, cough, shortness of breath

Patients (n=99)	
Signs and symptoms at admission	
Fever	82 (83%)
Cough	81 (82%)
Shortness of breath	31 (31%)
Muscle ache	11 (11%)
Confusion	9 (9%)
Headache	8 (8%)
Sore throat	5 (5%)
Rhinorrhoea	4 (4%)
Chest pain	2 (2%)
Diarrhoea	2 (2%)
Nausea and vomiting	1 (1%)
More than one sign or symptom	89 (90%)
Fever, cough, and shortness of breath	15 (15%)

Clinical Characteristics of Coronavirus Disease 2019 in China

W. Guan, Z. Ni, Yu Hu, W. Liang, C. Ou, J. He, L. Liu, H. Shan, C. Lei, D.S.C. Hui, B. Du, L. Li, G. Zeng, K.-Y. Yuen, R. Chen, C. Tang, T. Wang, P. Chen, J. Xiang,

- ▶ NEJM, February 28, 2020
- ▶ 1099 patients – 93.6% hospitalized
- ▶ Less than half of patients presented with fever
- ▶ Vast majority did develop fever during hospitalization
- ▶ Wide range of other reported symptoms
- ▶ Cough, sputum, and fatigue most common

Characteristic	All Patients (N = 1099)
Fever on admission	
Patients — no./total no. (%)	473/1081 (43.8)
Fever during hospitalization	
Patients — no./total no. (%)	975/1099 (88.7)
Symptoms — no. (%)	
Conjunctival congestion	9 (0.8)
Nasal congestion	53 (4.8)
Headache	150 (13.6)
→ Cough	745 (67.8)
Sore throat	153 (13.9)
→ Sputum production	370 (33.7)
→ Fatigue	419 (38.1)
Hemoptysis	10 (0.9)
Shortness of breath	205 (18.7)
Nausea or vomiting	55 (5.0)
Diarrhea	42 (3.8)
Myalgia or arthralgia	164 (14.9)
Chills	126 (11.5)

Clinical Characteristics of Coronavirus Disease 2019 in China

► 15.7% of 1099 patients had “severe disease”

Table 3. Complications, Treatments, and Clinical Outcomes.

Variable	All Patients (N = 1099)	Disease Severity	
		Nonsevere (N = 926)	Severe (N = 173)
Complications			
Septic shock — no. (%)	12 (1.1)	1 (0.1)	11 (6.4)
Acute respiratory distress syndrome — no. (%)	37 (3.4)	10 (1.1)	27 (15.6)
Acute kidney injury — no. (%)	6 (0.5)	1 (0.1)	5 (2.9)
Disseminated intravascular coagulation — no. (%)	1 (0.1)	0	1 (0.6)
Rhabdomyolysis — no. (%)	2 (0.2)	2 (0.2)	0
Physician-diagnosed pneumonia — no./total no. (%)	972/1067 (91.1)	800/894 (89.5)	172/173 (99.4)
Median time until development of pneumonia (IQR) — days*			
After initial Covid-19 diagnosis	0.0 (0.0–1.0)	0.0 (0.0–1.0)	0.0 (0.0–2.0)
After onset of Covid-19 symptoms	3.0 (1.0–6.0)	3.0 (1.0–6.0)	5.0 (2.0–7.0)

Clinical Characteristics of Coronavirus Disease 2019 in China

W. Guan, Z. Ni, Yu Hu, W. Liang, C. Ou, J. He, L. Liu, H. Shan, C. Lei, D.S.C. Hui, B. Du, L. Li, G. Zeng, K.-Y. Yuen, R. Chen, C. Tang, T. Wang, P. Chen, J. Xiang,

► 6.1% experienced the primary composite endpoint:

- ICU admission - 5%
- invasive MV - 2.3%
- death - 1.4%

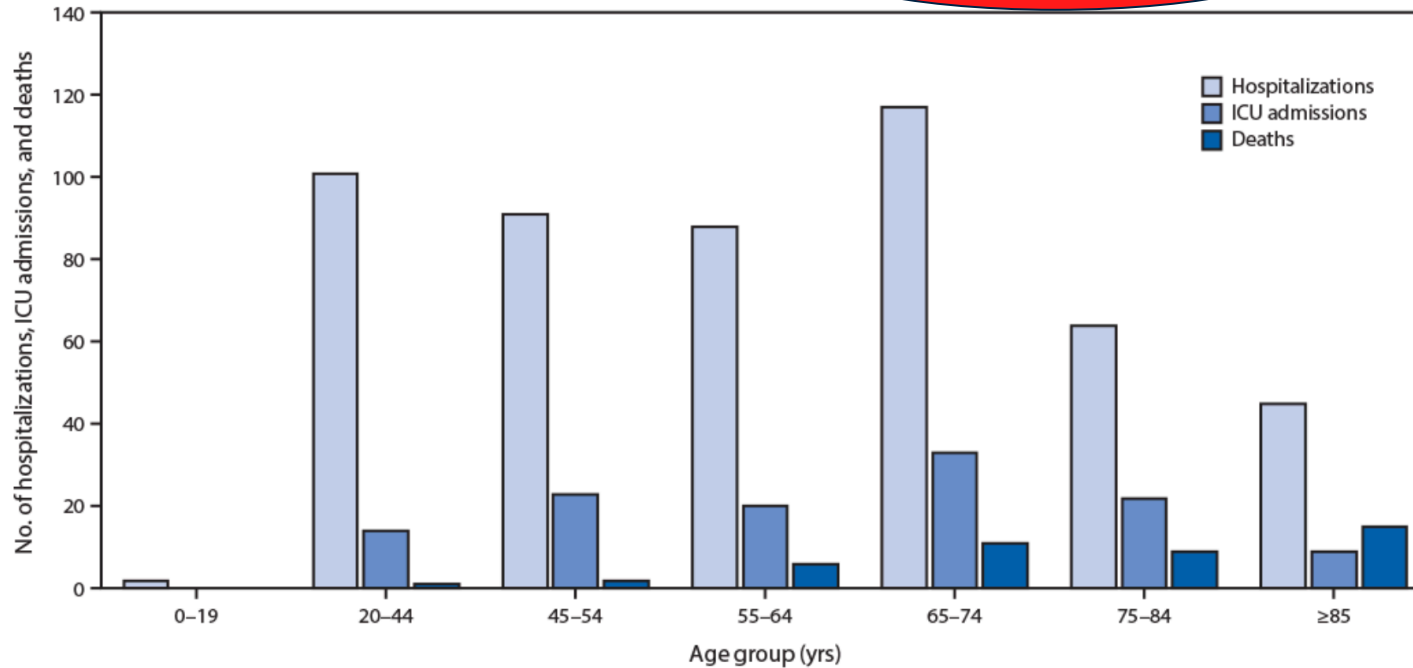
► Characteristics associated with worst outcomes

- Older age
- Comorbid illness

Characteristic	Presence of Primary Composite End Point†	
	Yes (N = 67)	No (N = 1032)
Age		
Median (IQR) — yr	63.0 (53.0–71.0)	46.0 (35.0–57.0)
Distribution — no./total no. (%)		
0–14 yr	0	9/946 (1.0)
15–49 yr	12/65 (18.5)	545/946 (57.6)
50–64 yr	21/65 (32.3)	271/946 (28.6)
≥65 yr	32/65 (49.2)	121/946 (12.8)
Coexisting disorder — no. (%)		
Any	39 (58.2)	222 (21.5)
Chronic obstructive pulmonary disease	7 (10.4)	5 (0.5)
Diabetes	18 (26.9)	63 (6.1)
Hypertension	24 (35.8)	141 (13.7)
Coronary heart disease	6 (9.0)	21 (2.0)
Cerebrovascular disease	4 (6.0)	11 (1.1)
Hepatitis B infection¶	1 (1.5)	22 (2.1)
Cancer	1 (1.5)	9 (0.9)
Chronic renal disease	2 (3.0)	6 (0.6)
Immunodeficiency	0	2 (0.2)

US Disease Outcomes – CDC.gov

FIGURE 2. Coronavirus disease 2019 (COVID-19) hospitalizations,* intensive care unit (ICU) admissions,[†] and deaths,[§] by age group — United States, February 12– March 16, 2020



* Hospitalization status missing or unknown for 1,514 cases.

[†] ICU status missing or unknown for 2,253 cases.

[§] Illness outcome or death missing or unknown for 2,001 cases.

- ▶ 4,226 total US cases at that time
- ▶ Highest fatality in age ≥ 85 (10-27%)
- ▶ Overall in US, persons age ≥ 65 :
 - 31% of cases
 - 45% of hospitalizations
 - 53% of ICU admissions
 - 80% of deaths
- ▶ Of ICU admissions
 - 36% age 45-64
 - 12% age 20-44

NYC and Philadelphia

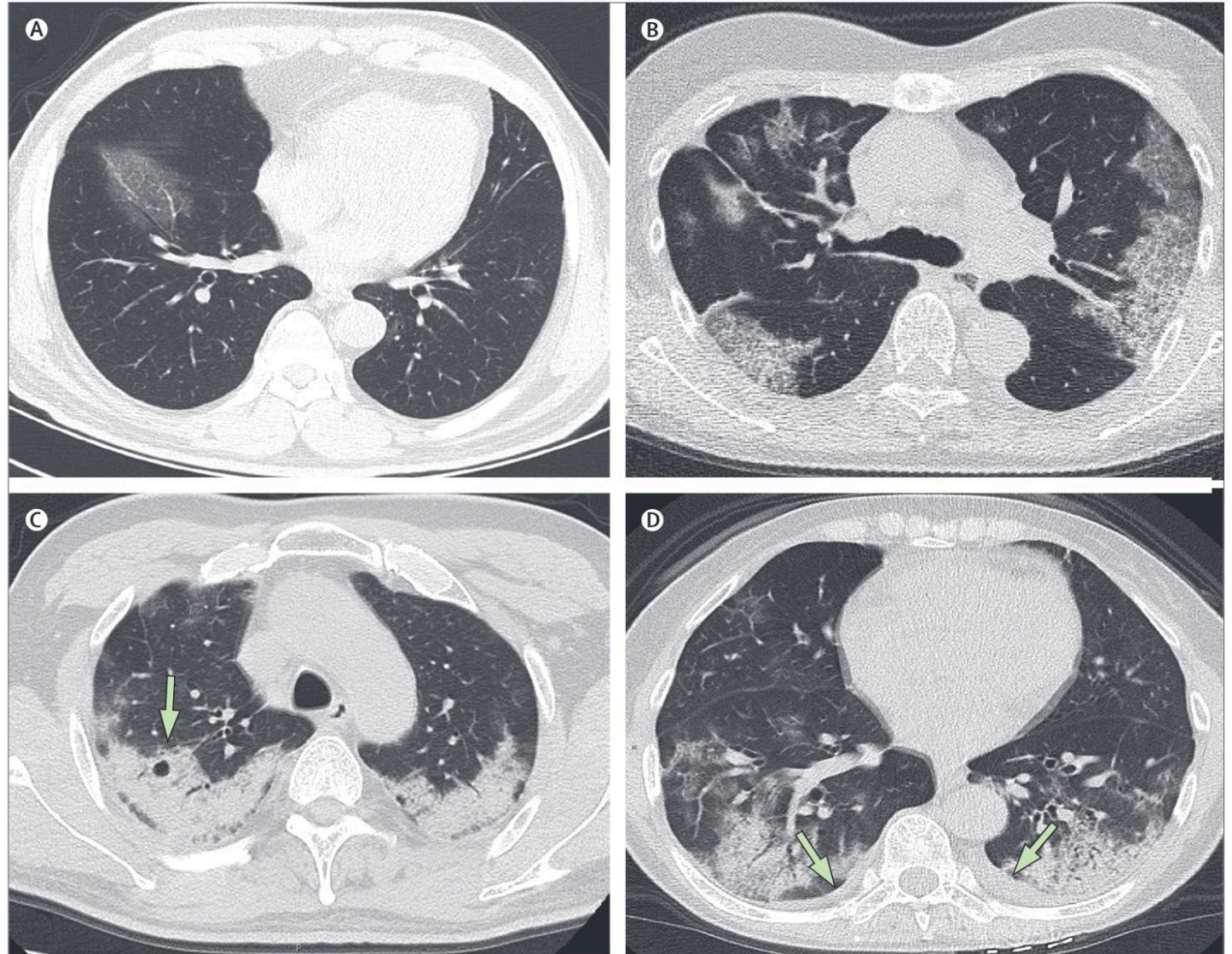
- ▶ In Philadelphia, half of cases <40 years
- ▶ In NYC, majority of cases <65 years

NYC COVID-19 Cases

Total Cases	
Total	32308
Median Age (Range)	48 (0-105)
Age Group	
- 0 to 17	611 (2%)
- 18 to 44	13794 (43%)
- 45 to 64	11146 (35%)
- 65 to 74	3790 (12%)
- 75 and over	2897 (9%)
- Unknown	70
Age 50 and over	
- Yes	15074 (47%)
- No	17164 (53%)
Sex	
- Female	14293 (44%)
- Male	17971 (56%)
- Unknown	44
Borough	
- Bronx	6145 (19%)
- Brooklyn	8451 (26%)
- Manhattan	5438 (17%)
- Queens	10373 (32%)
- Staten Island	1866 (6%)
- Unknown	35
Deaths	678

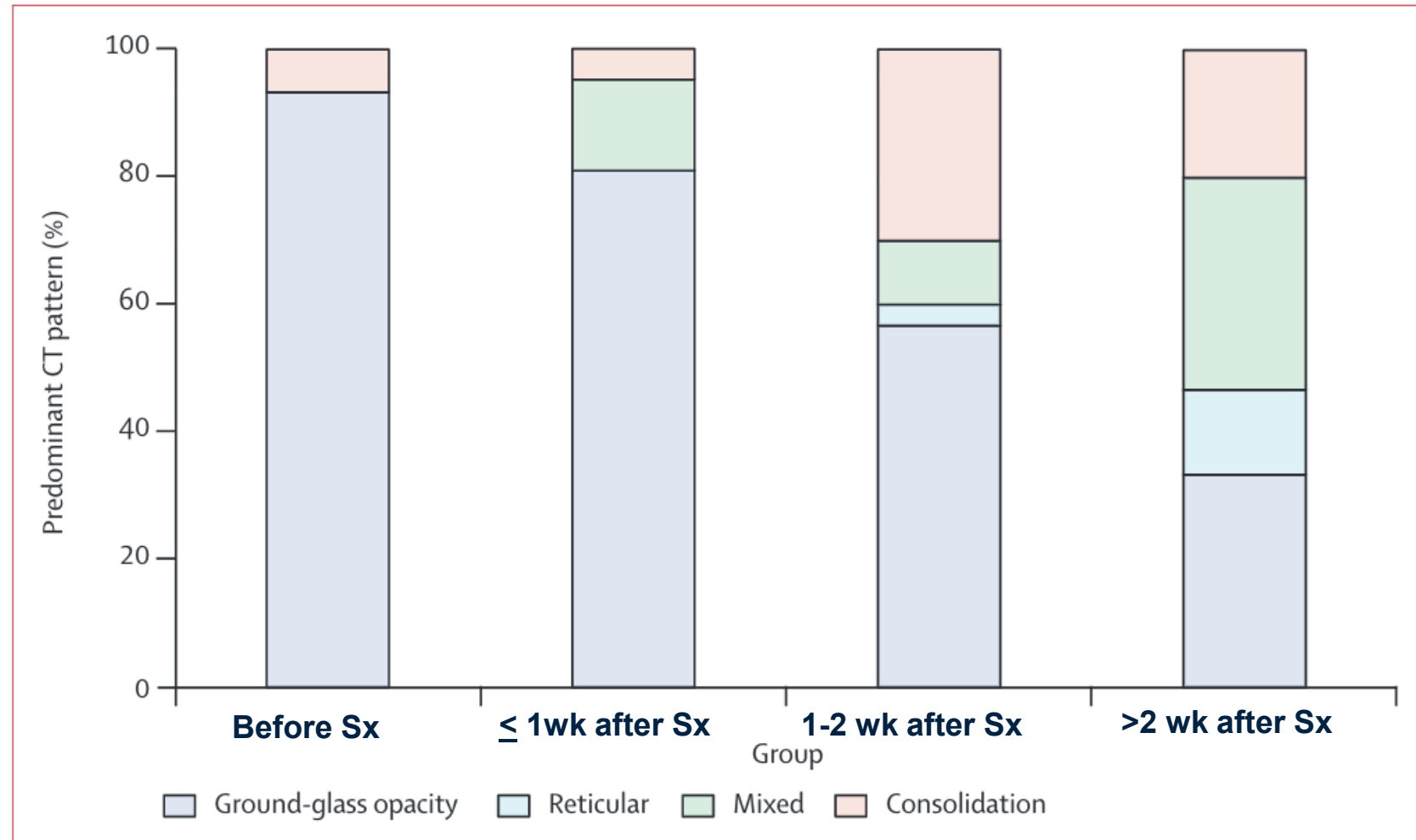
Imaging Findings

- ▶ A: 56M, d3 after Sx onset
- ▶ B: 74W, d10 after Sx onset
- ▶ C: 61W, d20 after Sx onset
- ▶ D: 63W, d17 after Sx onset

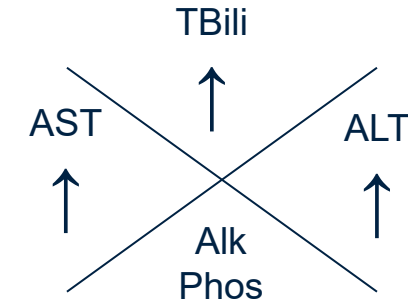
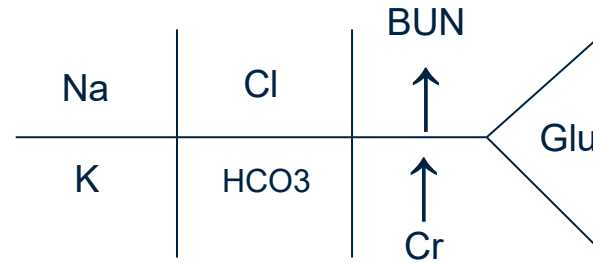
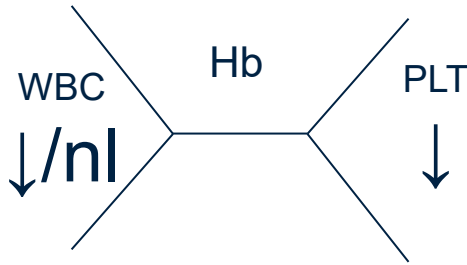


Imaging Findings

- Distribution CT patterns at various timepoints from symptom onset

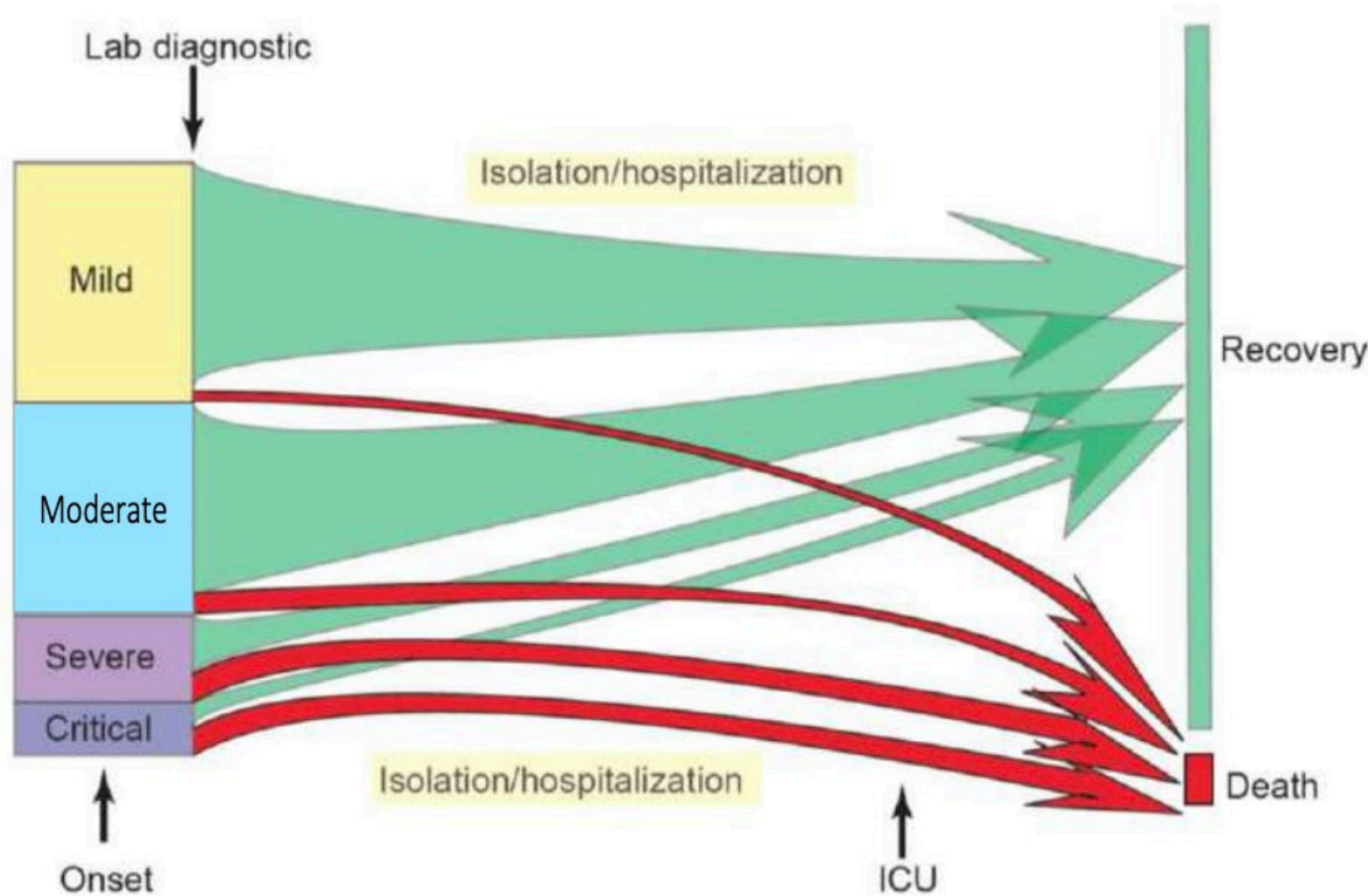


Laboratory Findings



- ▶ Leukopenia and lymphopenia (80%+)
- ▶ IL-6, Ferritin elevated
- ▶ D-Dimer, CRP, LDH elevated
- ▶ Procalcitonin generally low → may be high with bacterial superinfection

Disease Progression



Median time from onset to recovery:

- ▶ Mild: 2 weeks
- ▶ Severe: 3-6 weeks

Onset to development of severe disease:

- ▶ 1 week

Among those who have died, time of symptom onset to death:

- ▶ 2-8 weeks

Decompensation – Mainly Anecdotal

▶ Respiratory Failure

- Rapidly progressive from hospital admission (often 7-10d out from symptom onset)

▶ Shock

- Onset described when respiratory failure seems to be resolving
- Described as cold/clamped, POCUS demonstrating impaired cardiac function

▶ Cardiac Arrest

- VT/VF
- PEA

Washington ICU Outcomes

- ▶ Case series of 21 critically ill patients from Washington State
- ▶ Published 3/19/20
- ▶ Over half had severe ARDS
- ▶ 2/3 developed shock requiring vasopressors
- ▶ 19% AKI
- ▶ 1/3 developed a cardiomyopathy

Table 2. Clinical Measures During the Course of Illness and Outcomes of 21 Critically Ill Patients With Coronavirus Disease 2019

	No. (%) of patients ^a
Clinical measures	
Acute respiratory distress syndrome (ARDS) ^b	
None	1 (4.8)
Mild	2 (9.5)
Moderate	6 (28.6)
Severe	12 (57.1)
Required mechanical ventilation	15 (71.0)
Use of vasopressors	14 (67.0)
Acute kidney failure ^d	4 (19.1)
Cardiomyopathy ^e	7 (33.3)
Acute hepatic injury ^f	3 (14.3)
Seizures	1 (4.8)
Length of follow-up, mean (range), d	5.2 (1-10)
Outcomes	
Died	11 (52.4)
Survived to transfer out of ICU	2 (9.5)
Remains critically ill and requires mechanical ventilation	8 (38.1)
Length of follow-up for those who survived or remain critically ill, mean (range), d	7.5 (5-10)



Critical Care Management

Roger Kim



Outline

- ▶ Respiratory failure
- ▶ Mechanical ventilation
- ▶ Hemodynamic management
- ▶ Novel strategies / future directions

Respiratory failure algorithms

UPHS guidelines

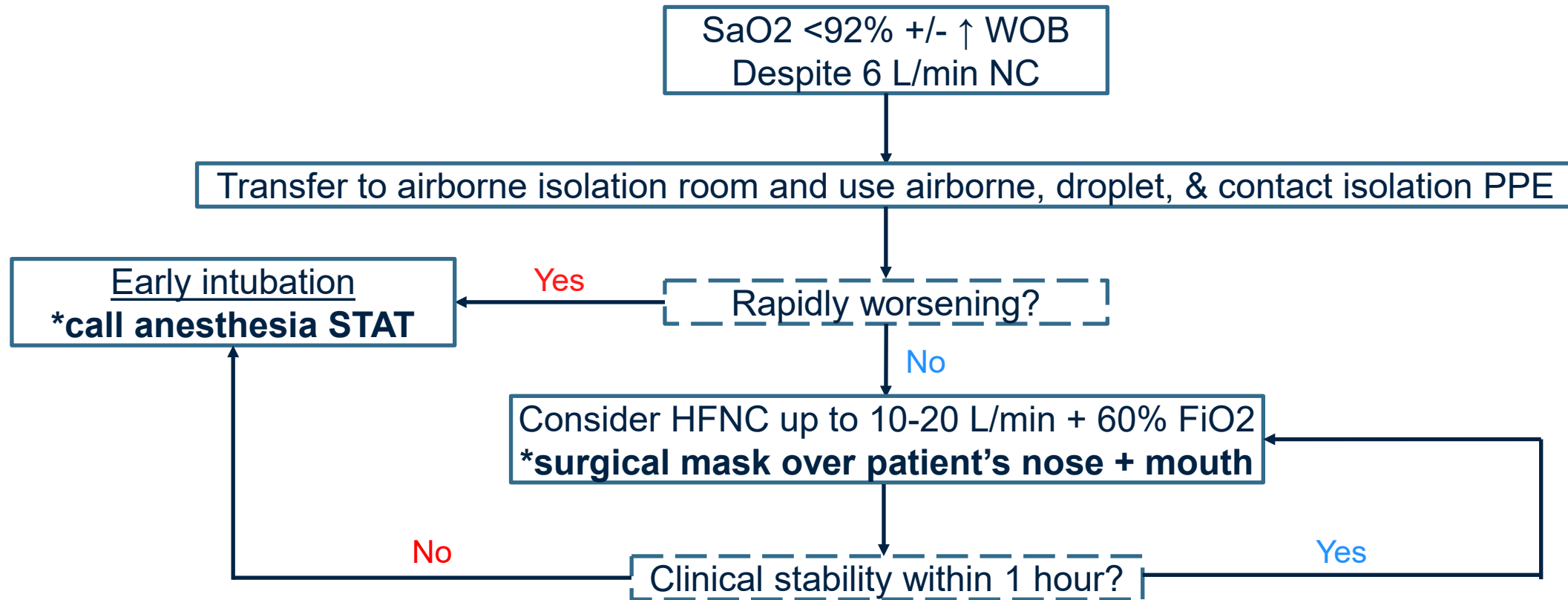
March 27, 2020



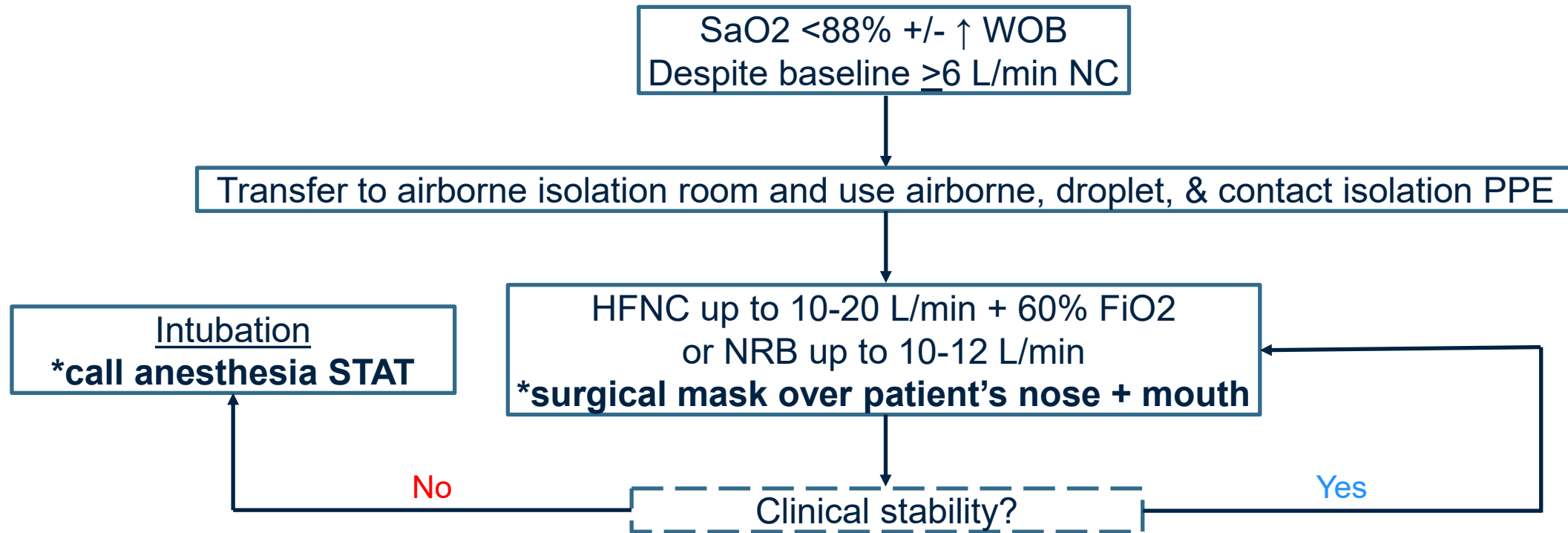
Respiratory support general principles

- ▶ Favor early intubation
- ▶ Avoid NIPPV (i.e. CPAP, BiPAP)
- ▶ Limit HFNC use to 10-20 L/min flow rate and 60% FiO₂

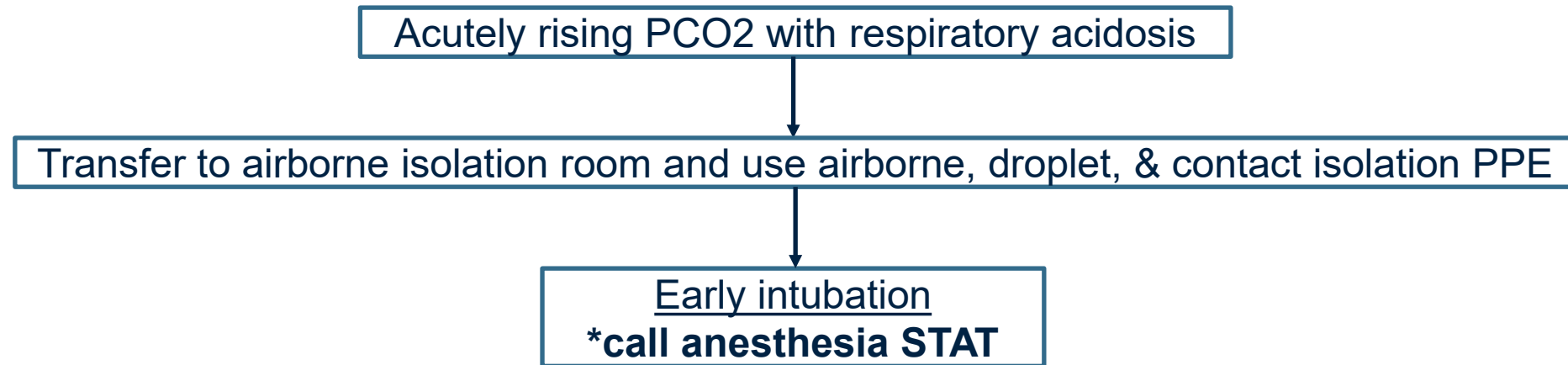
Acute hypoxic respiratory failure



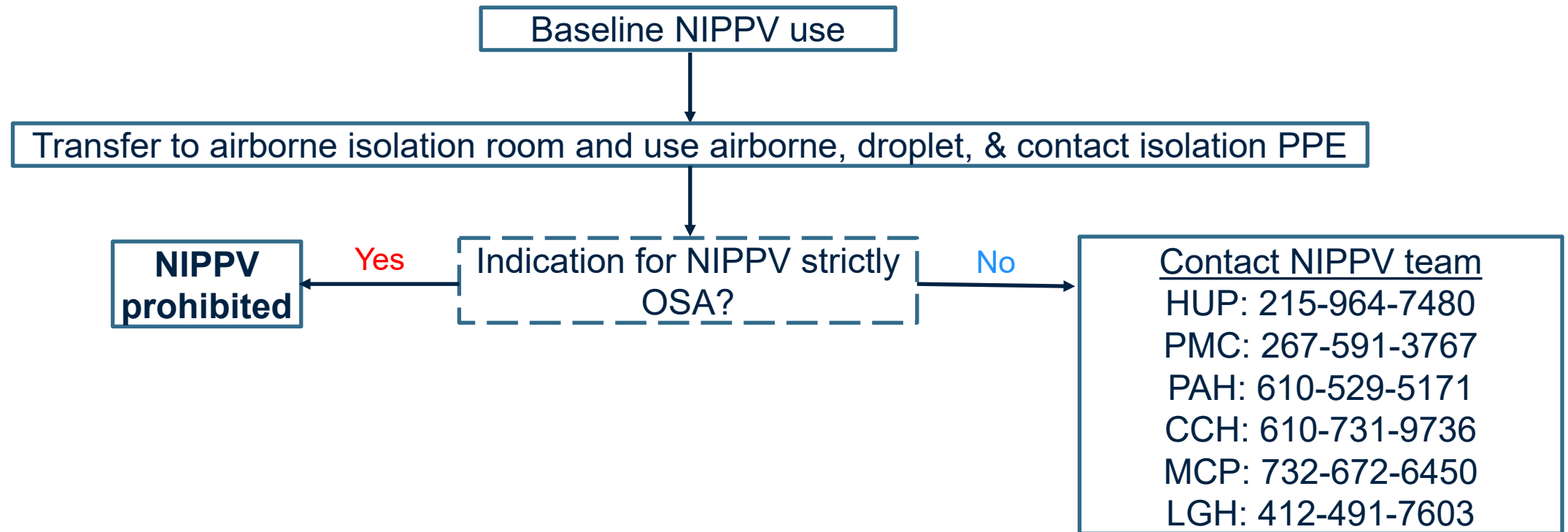
Acute on chronic hypoxic respiratory failure



Acute / acute on chronic hypercapnic respiratory failure



Stable chronic hypercapnic respiratory failure



ARDS mechanical ventilation strategies

UPHS guidelines
March 27, 2020

SCCM COVID-19 guidelines
March 2020



ARDS mechanical ventilation general principles

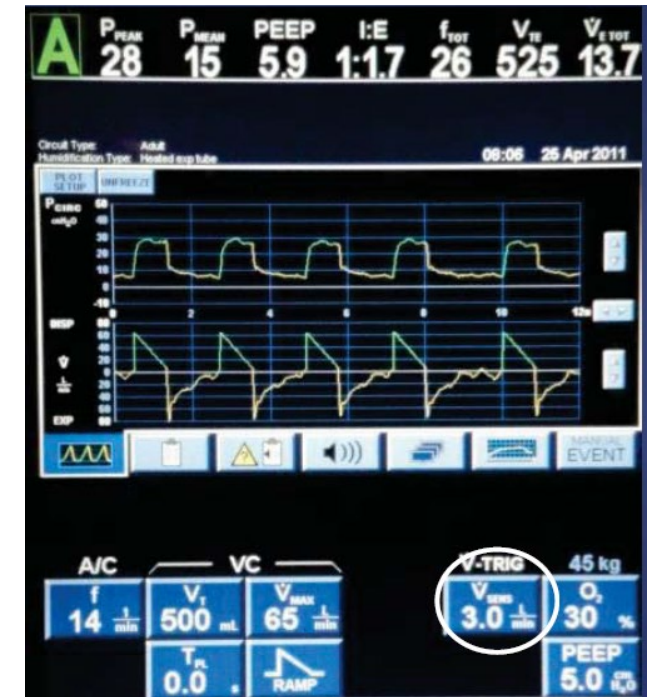
- ▶ Low-stretch (lung protective) ventilation
- ▶ High PEEP strategy
- ▶ Conservative fluid strategy
- ▶ Refractory hypoxemia
 - Neuromuscular blocking agents (NMBA)
 - Prone ventilation

ARDS mechanical ventilation general principles

- ▶ **Low-stretch (lung protective) ventilation**
- ▶ High PEEP strategy
- ▶ Conservative fluid strategy
- ▶ Refractory hypoxemia
 - Neuromuscular blocking agents (NMBA)
 - Prone ventilation

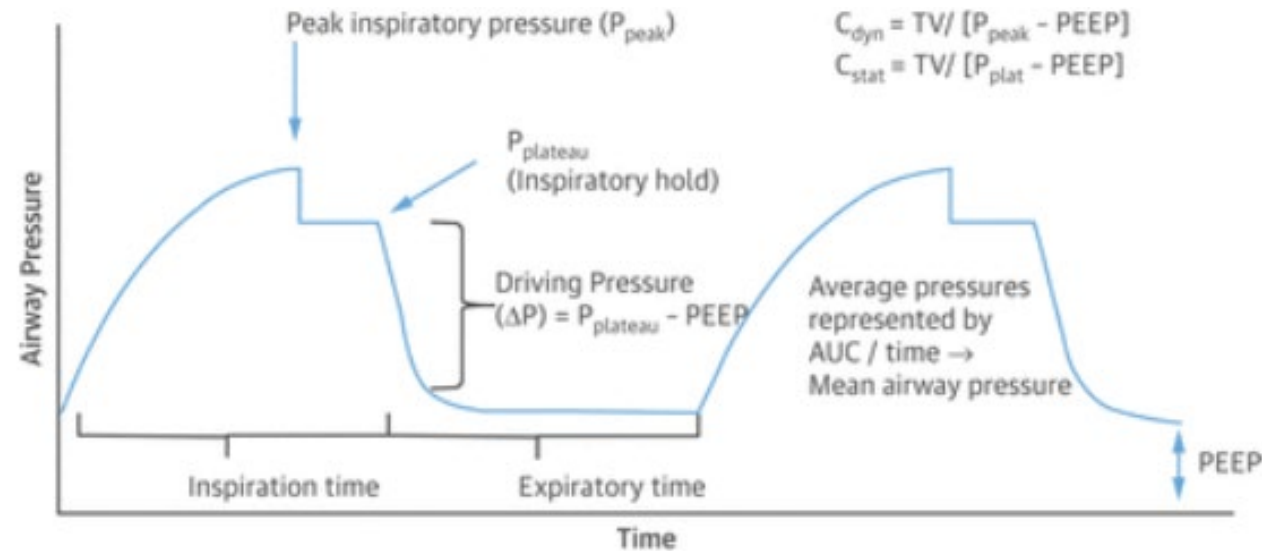
Low-stretch (lung protective) ventilation

- ▶ Volume assist control (VAC) with low tidal volumes
 - **Vt 4-8 mL/kg of predicted ideal body weight (start at 6 mL/kg)**
- ▶ Target plateau pressure (P_{plat}) < 30 cm H₂O
- ▶ Goal pH: 7.30-7.45
- ▶ Goal SpO₂: 92-96%
- ▶ Ventilator dyssynchrony is common
 - Adequate sedation is required
 - Consider RASS goal of -2 to -3



High PEEP strategy

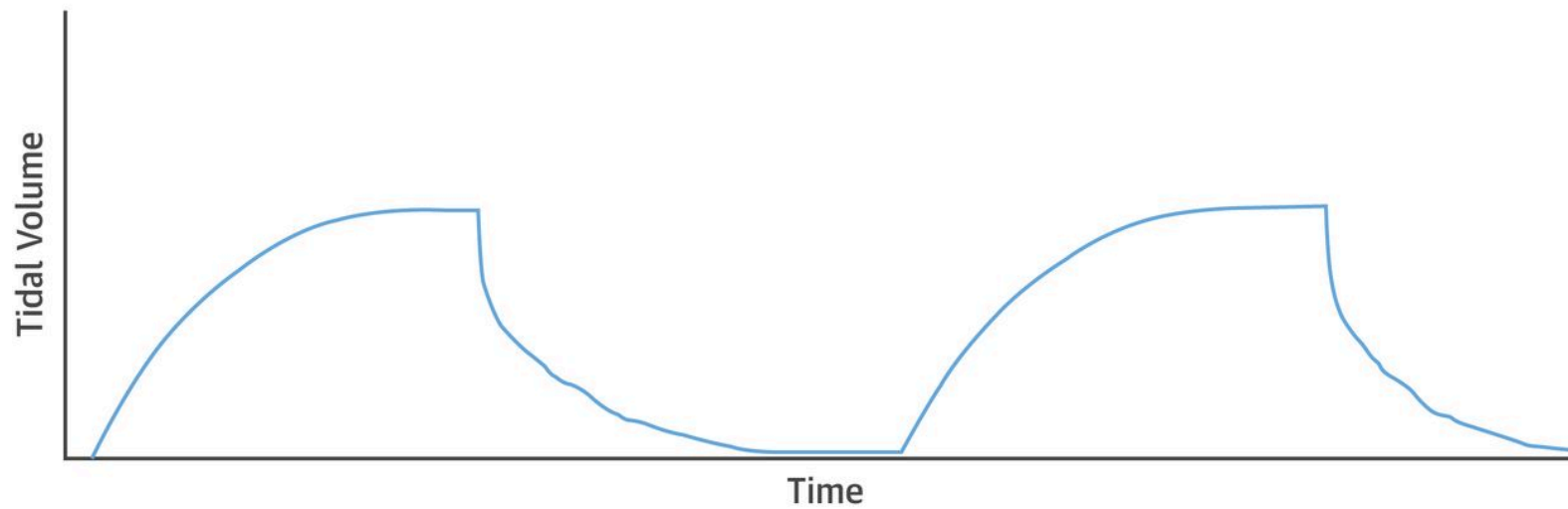
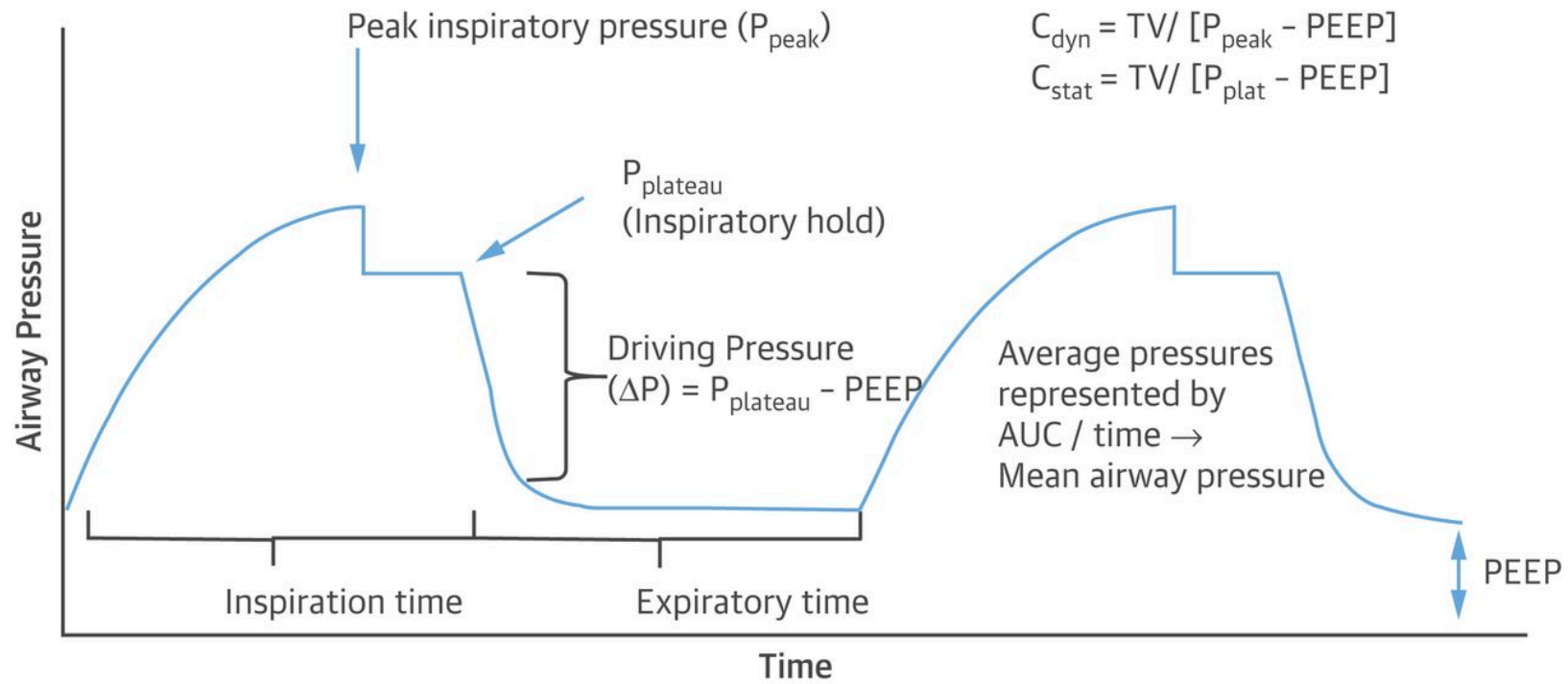
- ▶ PEEP-responsive ARDS with driving pressures <15 cm H₂O consistently reported
- ▶ Start with PEEP of ≥14-18 cm H₂O
- ▶ Risk of PTX and hemodynamic compromise



Higher PEEP/lower FiO₂

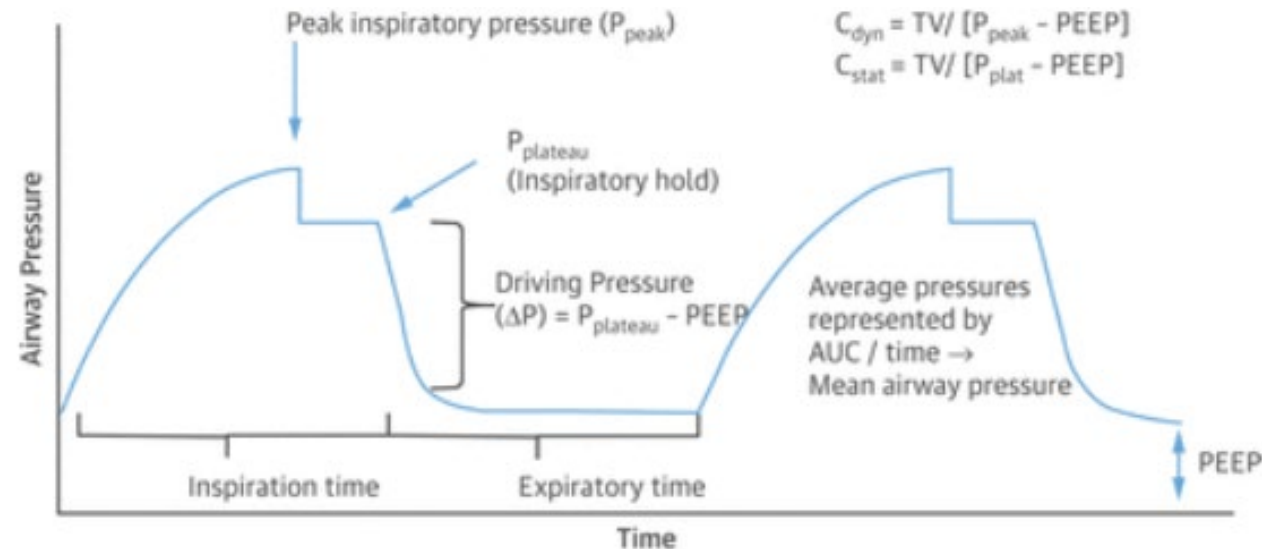
FiO₂	0.3	0.3	0.3	0.3	0.3	0.4	0.4	0.5
PEEP	5	8	10	12	14	14	16	16

FiO₂	0.5	0.5-0.8	0.8	0.9	1.0	1.0
PEEP	18	20	22	22	22	24



High PEEP strategy

- ▶ PEEP-responsive ARDS with driving pressures <15 cm H₂O consistently reported
- ▶ Start with PEEP of ≥14-18 cm H₂O
- ▶ Risk of PTX and hemodynamic compromise



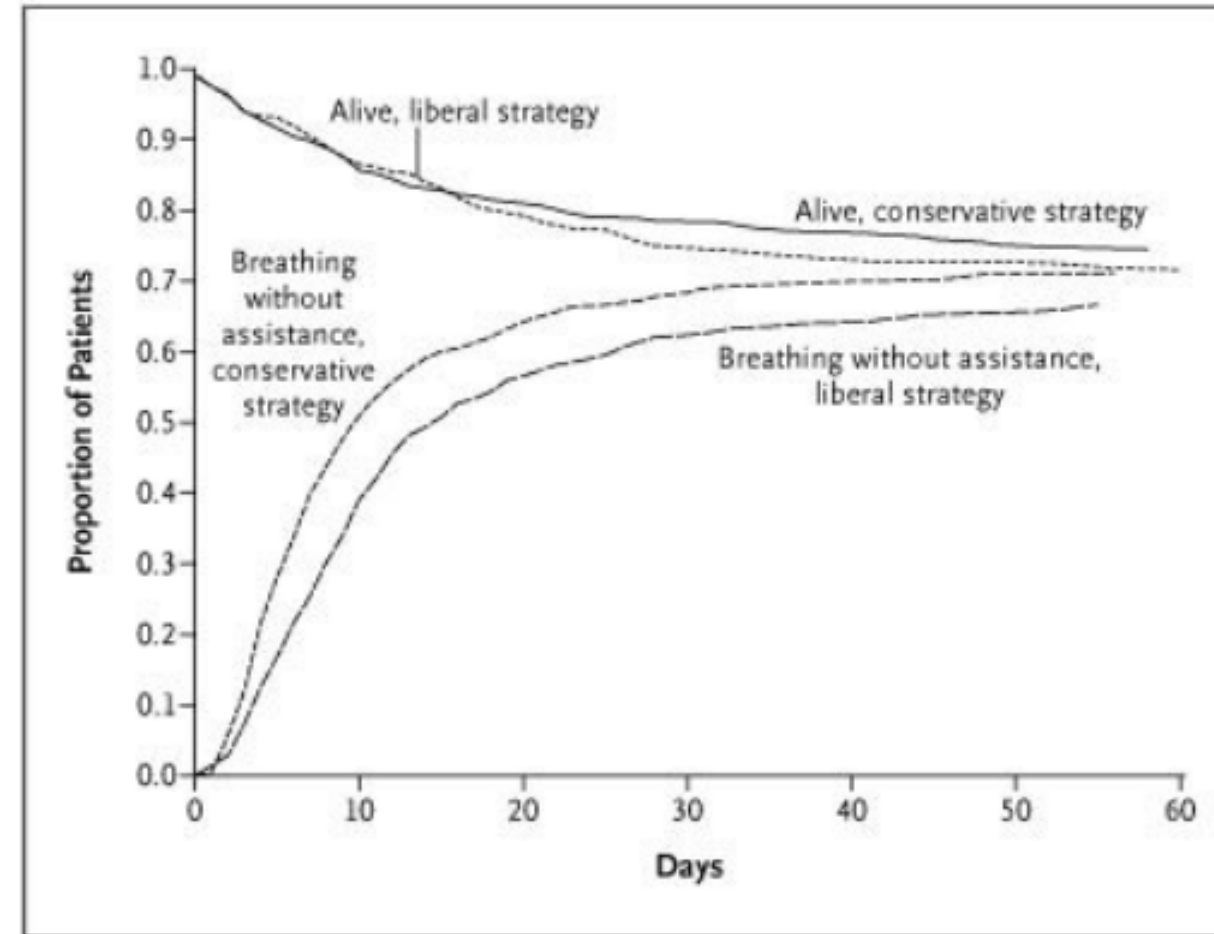
Higher PEEP/lower FiO₂

FiO₂	0.3	0.3	0.3	0.3	0.3	0.4	0.4	0.5
PEEP	5	8	10	12	14	14	16	16

FiO₂	0.5	0.5-0.8	0.8	0.9	1.0	1.0
PEEP	18	20	22	22	22	24

Conservative fluid strategy

- ▶ No consensus on definition
- ▶ Diurese to avoid obvious volume overload
- ▶ Diurese if SpO₂ <92% despite optimization of ventilator mechanics
- ▶ Consider diuresis if on low-dose pressors with normal renal function



Refractory hypoxemia – recommended strategies

Neuromuscular blocking agents (NMBA)

- ▶ Intermittent boluses to facilitate lung protective ventilation
- ▶ Continuous infusion ≤ 48 hrs for prone ventilation, persistent $P_{\text{plat}} > 30$ cm H₂O, or persistent ventilator dyssynchrony

Prone ventilation

- ▶ Use if $P:F \leq 150$, $FiO_2 \geq 60\%$, and $PEEP \geq 5$ cm H₂O after ≥ 12 hours of ventilator support
- ▶ Prone 16-18 hours per day
- ▶ Stop:
 - When $P:F > 150$ with $PEEP \leq 10$ cm H₂O and $FiO_2 \leq 60\%$ in supine position for ≥ 4 hours
 - If prone position decreases $P:F$ by $> 20\%$ compared to supine position

Refractory hypoxemia – strategies to consider

Traditional recruitment maneuvers

- ▶ 30-40 cm H₂O for 30-40 s
- ▶ If oxygenation improves, use higher PEEP

Bronchoscopy

ONLY in following situations:

- ▶ Complete lung atelectasis from mucous plugging with worsening hypoxemia
- ▶ Massive hemoptysis with need to clear blood/clot and place bronchial blocker
- ▶ Unable to obtain tracheal aspirate for VAP workup

VV-ECMO

Consider if all other interventions exhausted AND any 1 of following:

- ▶ Injurious ventilator settings necessary to achieve adequate oxygenation
- ▶ Uncontrolled respiratory acidosis
- ▶ Right heart failure with persistent organ dysfunction despite lung protective ventilation
 - May need to switch to VA-ECMO if does not improve with VV-ECMO

Refractory hypoxemia – **NOT** recommended strategies

Inhaled epoprostenol (i.e. Flolan, Veletri)

- ▶ No mortality benefit
- ▶ May clog vent filter and increase risk of aerosolization
- ▶ *Can consider a trial of inhaled nitric oxide after discussion with pharmacy*

Staircase (incremental PEEP) recruitment maneuvers

- ▶ Defined as incremental increases in PEEP from 25 to 35 to 45 cm H₂O for 1-2 min. each
- ▶ May be associated with increased mortality

Hemodynamic management

UPHS guidelines
March 27, 2020

SCCM COVID-19 guidelines
March 2020



Septic shock management

- ▶ Empiric antibiotics in mechanically ventilated patients
- ▶ Conservative isotonic crystalloid fluids (LR > NS) for acute resuscitation
 - Preferred over hydroxyethyl starches, dextrans, gelatins, or albumin
- ▶ Norepinephrine = preferred 1st line vasoactive agent
 - If not available, consider vasopressin or epinephrine
 - Preferred over dopamine
- ▶ Vasopressin = preferred 2nd line vasoactive agent
 - If not available, consider epinephrine
- ▶ MAP goal: 60-65 mmHg
- ▶ Consider “stress-dose” steroids (hydrocortisone 200 mg/d) for refractory shock

COVID-19 hemodynamic considerations

- ▶ Start norepinephrine at 0.05-0.1 mcg/kg/min immediately after intubation and titrate accordingly
- ▶ Presumed viral myocarditis → cardiac dysfunction / fluid overload
 - Favor negative fluid balance without causing organ hypoperfusion
 - Consider diuresis if POCUS reveals non-collapsible IVC
 - Dynamic hemodynamic reassessment with POCUS TTE
 - Consider VA-ECMO for severe myocarditis causing cardiogenic shock
- ▶ Ensure adequate preload in setting of high PEEP ventilatory strategy
 - Consider careful IVF boluses if PEEP >15cm H₂O

Novel strategies / future directions



Helmet NIPPV



Helmet interface
for noninvasive ventilation

Awake prone ventilation

- ▶ Floor patients with hypoxic respiratory failure on supplemental O₂ (including HFNC)
- ▶ **NOT** recommended in patients with:
 - Chronic lung disease
 - Chest tubes
 - Spinal instability
 - Cardiogenic pulmonary edema
 - GCS <15
 - PaCO₂ >45 mmHg
- ▶ Prone for ≥ 2 -4 hours bid as tolerated



BMI-based PEEP ventilatory strategy

BMI	Starting PEEP
<35	10
35-50	12
>50	15

Future directions

- ▶ Define PEEP weaning and prolonged SBT protocol
- ▶ Standardize approach to cardiac monitoring
 - Daily EKG not recommended
 - Early POCUS vs. TTE?
- ▶ Standardize sedation protocol
 - Concerns about propofol (↑TG's & LFT's)
 - Cisatracurium shortage
- ▶ Standardize laboratory testing frequency
 - D-dimer, procalcitonin, LFT's, LDH, ferritin, WBC w/ diff, CRP, IL-6, coags

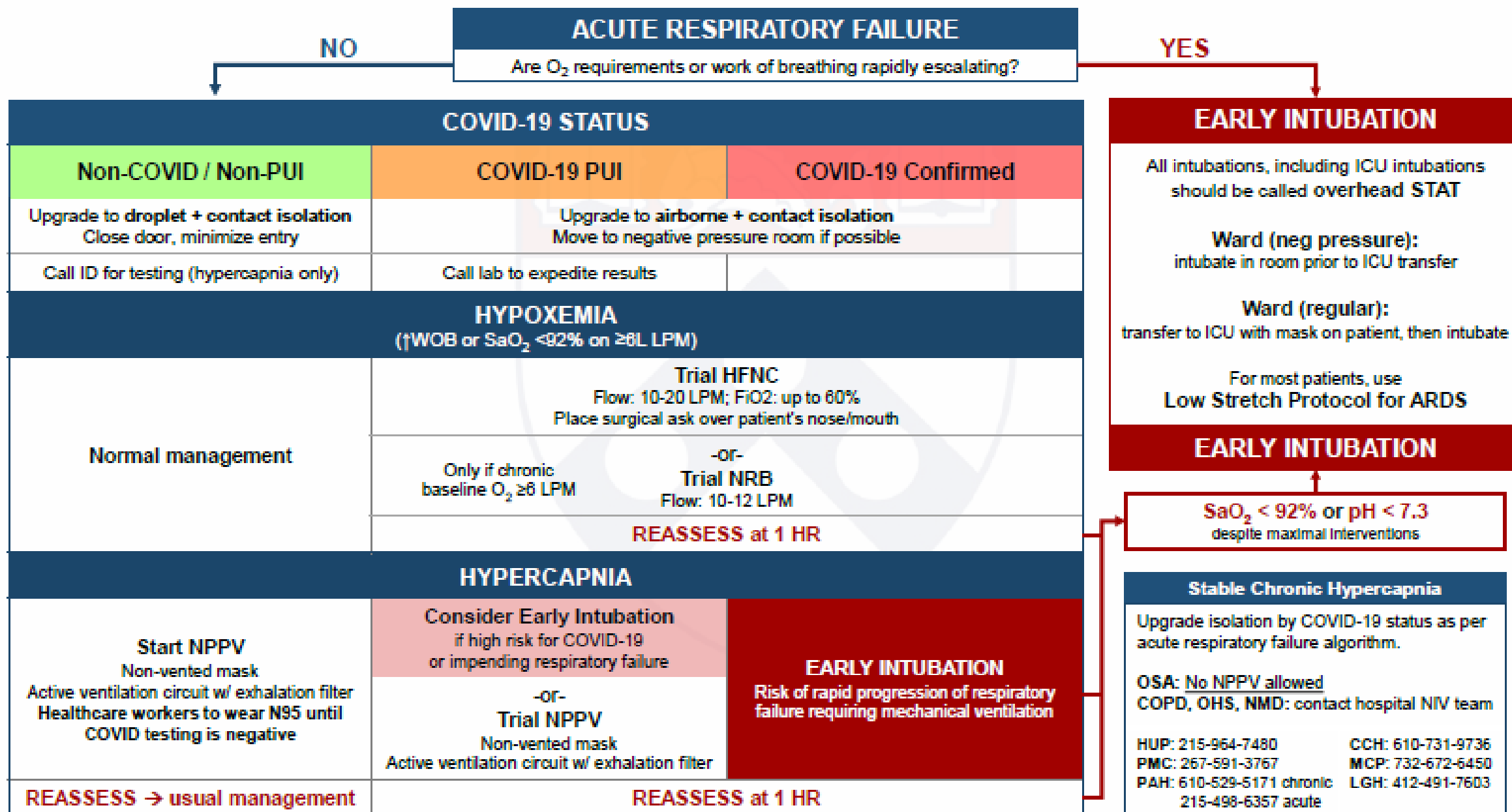


Supplemental slides / figures



Penn Medicine Tip Sheet: Escalation of Care for Respiratory Failure

Updated 3/28/2020 – [newest version here](#)



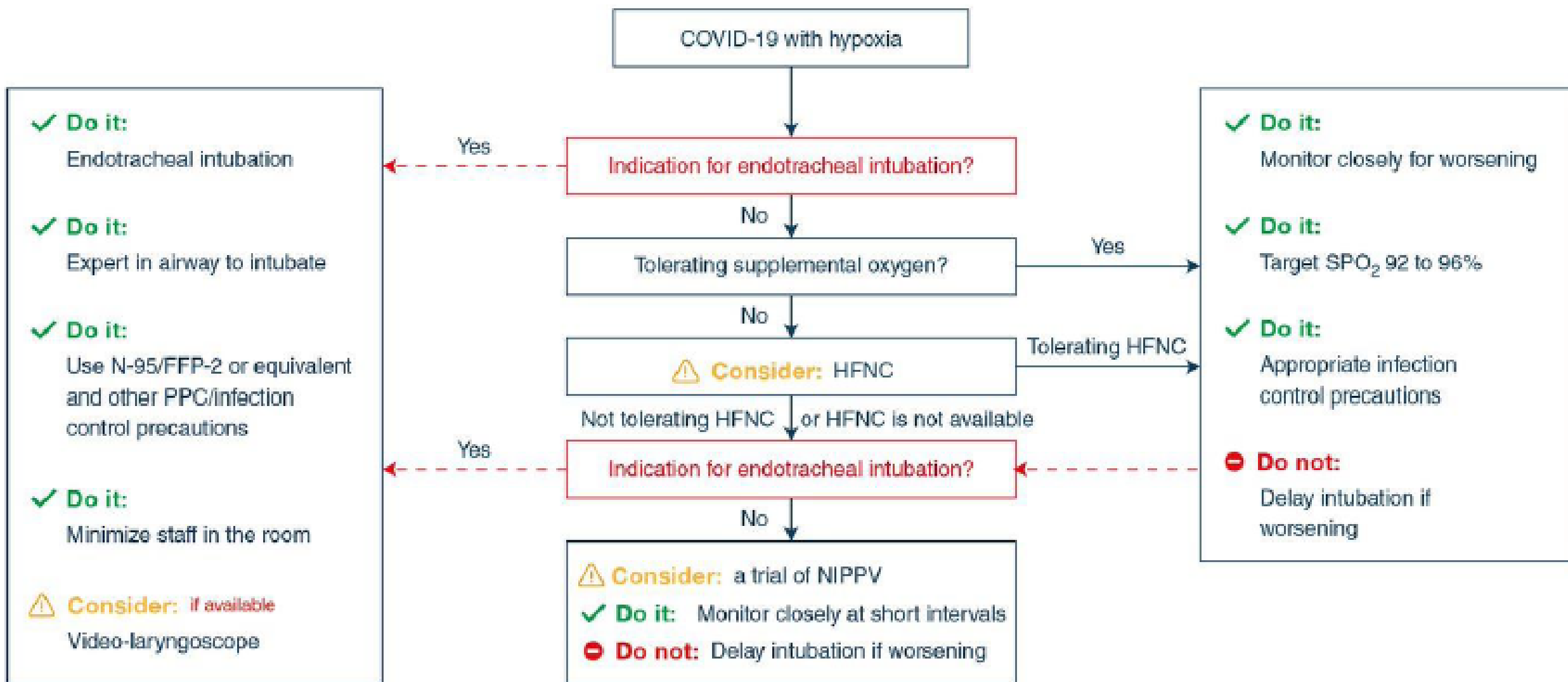


Table 3. The Berlin Definition of Acute Respiratory Distress Syndrome

Acute Respiratory Distress Syndrome	
Timing	Within 1 week of a known clinical insult or new or worsening respiratory symptoms
Chest imaging ^a	Bilateral opacities—not fully explained by effusions, lobar/lung collapse, or nodules
Origin of edema	Respiratory failure not fully explained by cardiac failure or fluid overload Need objective assessment (eg, echocardiography) to exclude hydrostatic edema if no risk factor present
Oxygenation ^b	
Mild	$200 \text{ mm Hg} < \text{PaO}_2/\text{FiO}_2 \leq 300 \text{ mm Hg}$ with PEEP or CPAP $\geq 5 \text{ cm H}_2\text{O}$ ^c
Moderate	$100 \text{ mm Hg} < \text{PaO}_2/\text{FiO}_2 \leq 200 \text{ mm Hg}$ with PEEP $\geq 5 \text{ cm H}_2\text{O}$
Severe	$\text{PaO}_2/\text{FiO}_2 \leq 100 \text{ mm Hg}$ with PEEP $\geq 5 \text{ cm H}_2\text{O}$

Abbreviations: CPAP, continuous positive airway pressure; FiO_2 , fraction of inspired oxygen; PaO_2 , partial pressure of arterial oxygen; PEEP, positive end-expiratory pressure.

^aChest radiograph or computed tomography scan.

^bIf altitude is higher than 1000 m, the correction factor should be calculated as follows: $[\text{PaO}_2/\text{FiO}_2 \times (\text{barometric pressure}/760)]$.

^cThis may be delivered noninvasively in the mild acute respiratory distress syndrome group.

NIH PREDICTED BODY WEIGHT (PBW) / TIDAL VOLUME CHART															
MALES								FEMALES							
HEIGHT		PBW	4	5	6	7	8	HEIGHT		PBW	4	5	6	7	8
Feet	Inches	Male	ml/kg	ml/kg	ml/kg	ml/kg	ml/kg	Feet	Inches	Female	ml/kg	ml/kg	ml/kg	ml/kg	ml/kg
4' 10"	58	45.4	180	230	270	320	360	4' 7"	55	34	140	170	200	240	270
4' 11"	59	47.7	190	240	290	330	380	4' 8"	56	36.3	150	180	220	250	290
5' 0"	60	50	200	250	300	350	400	4' 9"	57	38.6	150	190	230	270	310
5' 1"	61	52.3	210	260	310	370	420	4' 10"	58	40.9	160	200	250	290	330
5' 2"	62	54.6	220	270	330	380	440	4' 11"	59	43.2	170	220	260	300	350
5' 3"	63	56.9	230	280	340	400	460	5' 0"	60	45.5	180	230	270	320	360
5' 4"	64	59.2	240	300	360	410	470	5' 1"	61	47.8	190	240	290	330	380
5' 5"	65	61.5	250	310	370	430	490	5' 2"	62	50.1	200	250	300	350	400
5' 6"	66	63.8	260	320	380	450	510	5' 3"	63	52.4	210	260	310	370	420
5' 7"	67	66.1	260	330	400	460	530	5' 4"	64	54.7	220	270	330	380	440
5' 8"	68	68.4	270	340	410	480	550	5' 5"	65	57	230	290	340	400	460
5' 9"	69	70.7	280	350	420	490	570	5' 6"	66	59.3	240	300	360	420	470
5' 10"	70	73	290	370	440	510	580	5' 7"	67	61.6	250	310	370	430	490
5' 11"	71	75.3	300	380	450	530	600	5' 8"	68	63.9	260	320	380	450	510
6' 0"	72	77.6	310	390	470	540	620	5' 9"	69	66.2	260	330	400	460	530
6' 1"	73	79.9	320	400	480	560	640	5' 10"	70	68.5	270	340	410	480	550
6' 2"	74	82.2	330	410	490	580	660	5' 11"	71	70.8	280	350	420	500	570
6' 3"	75	84.5	340	420	510	590	680	6' 0"	72	73.1	290	370	440	510	580
6' 4"	76	86.8	350	430	520	610	690	6' 1"	73	75.4	300	380	450	530	600
6' 5"	77	89.1	360	450	530	620	710	6' 2"	74	77.7	310	390	470	540	620
6' 6"	78	91.4	370	460	550	640	730	6' 3"	75	80	320	400	480	560	640



NIH NHLBI ARDS Clinical Network Mechanical Ventilation Protocol Summary

INCLUSION CRITERIA: Acute onset of

1. $\text{PaO}_2/\text{FiO}_2 \leq 300$ (corrected for altitude)
2. Bilateral (patchy, diffuse, or homogeneous) infiltrates consistent with pulmonary edema
3. No clinical evidence of left atrial hypertension

PART I: VENTILATOR SETUP AND ADJUSTMENT

1. Calculate predicted body weight (PBW)
Males = $50 + 2.3 [\text{height (inches)} - 60]$
Females = $45.5 + 2.3 [\text{height (inches)} - 60]$
2. Select any ventilator mode
3. Set ventilator settings to achieve initial $V_T = 8 \text{ ml/kg PBW}$
4. Reduce V_T by 1 ml/kg at intervals ≤ 2 hours until $V_T = 6 \text{ ml/kg PBW}$.
5. Set initial rate to approximate baseline minute ventilation (not $> 35 \text{ bpm}$).
6. Adjust V_T and RR to achieve pH and plateau pressure goals below.

OXYGENATION GOAL: PaO_2 55-80 mmHg or SpO_2 88-95%

Use a minimum PEEP of 5 cm H_2O . Consider use of incremental FiO_2 /PEEP combinations such as shown below (not required) to achieve goal.

Lower PEEP/higher FiO_2

FiO_2	0.3	0.4	0.4	0.5	0.5	0.6	0.7	0.7
PEEP	5	5	8	8	10	10	10	12

FiO_2	0.7	0.8	0.9	0.9	0.9	1.0
PEEP	14	14	14	16	18	18-24

Higher PEEP/lower FiO_2

FiO_2	0.3	0.3	0.3	0.3	0.3	0.4	0.4	0.5
PEEP	5	8	10	12	14	14	16	16

FiO_2	0.5	0.5-0.8	0.8	0.9	1.0	1.0
PEEP	18	20	22	22	22	24

PLATEAU PRESSURE GOAL: $\leq 30 \text{ cm H}_2\text{O}$

Check P_{plat} (0.5 second inspiratory pause), at least q 4h and after each change in PEEP or V_T .

If $P_{\text{plat}} > 30 \text{ cm H}_2\text{O}$: decrease V_T by 1ml/kg steps (minimum = 4 ml/kg).

If $P_{\text{plat}} < 25 \text{ cm H}_2\text{O}$ and $V_T < 6 \text{ ml/kg}$, increase V_T by 1 ml/kg until $P_{\text{plat}} > 25 \text{ cm H}_2\text{O}$ or $V_T = 6 \text{ ml/kg}$.

If $P_{\text{plat}} < 30$ and breath stacking or dys-synchrony occurs: may increase V_T in 1ml/kg increments to 7 or 8 ml/kg if P_{plat} remains $\leq 30 \text{ cm H}_2\text{O}$.

pH GOAL: 7.30-7.45

Acidosis Management: (pH < 7.30)

If pH 7.15-7.30: Increase RR until pH > 7.30 or PaCO₂ < 25 (Maximum set RR = 35).

If pH < 7.15: Increase RR to 35.

If pH remains < 7.15, V_T may be increased in 1 ml/kg steps until pH > 7.15 (Pplat target of 30 may be exceeded).

May give NaHCO₃

Alkalosis Management: (pH > 7.45) Decrease vent rate if possible.

I: E RATIO GOAL: Recommend that duration of inspiration be ≤ duration of expiration.

PART II: WEANING

A. Conduct a SPONTANEOUS BREATHING TRIAL daily when:

1. FiO₂ ≤ 0.40 and PEEP ≤ 8 OR FiO₂ ≤ 0.50 and PEEP ≤ 5.
2. PEEP and FiO₂ ≤ values of previous day.
3. Patient has acceptable spontaneous breathing efforts. (May decrease vent rate by 50% for 5 minutes to detect effort.)
4. Systolic BP ≥ 90 mmHg without vasopressor support.
5. No neuromuscular blocking agents or blockade.

B. SPONTANEOUS BREATHING TRIAL (SBT):

If all above criteria are met and subject has been in the study for at least 12 hours, initiate a trial of UP TO 120 minutes of spontaneous breathing with FiO₂ ≤ 0.5 and PEEP ≤ 5:

1. Place on T-piece, trach collar, or CPAP ≤ 5 cm H₂O with PS ≤ 5
2. Assess for tolerance as below for up to two hours.
 - a. SpO₂ ≥ 90: and/or PaO₂ ≥ 60 mmHg
 - b. Spontaneous V_T ≥ 4 ml/kg PBW
 - c. RR ≤ 35/min
 - d. pH ≥ 7.3
 - e. No respiratory distress (distress= 2 or more)
 - HR > 120% of baseline
 - Marked accessory muscle use
 - Abdominal paradox
 - Diaphoresis
 - Marked dyspnea
3. If tolerated for at least 30 minutes, consider extubation.
4. If not tolerated resume pre-weaning settings.

Definition of UNASSISTED BREATHING (Different from the spontaneous breathing criteria as PS is not allowed)

1. Extubated with face mask, nasal prong oxygen, or room air, OR
2. T-tube breathing, OR
3. Tracheostomy mask breathing, OR
4. CPAP less than or equal to 5 cm H₂O **without pressure support or IMV assistance.**

COVID-19 with mild ARDS	COVID-19 with Mod to Severe ARDS	Rescue/Adjunctive therapy
✓ Do: Vt 4-8 ml/kg and $P_{plat} < 30$ cm H ₂ O	⚠ CONSIDER: Higher PEEP	? Uncertain: Antivirals, chloroquine, anti-IL6
✓ Do: Investigate for bacterial infection	⚠ CONSIDER: NMBA boluses to facilitate ventilation targets	⚠ CONSIDER: if proning, high P_{plat} , asynchrony NMBA infusion for 24 h
✓ Do: Target SPO2 92% - 96%	⚠ CONSIDER: if PEEP responsive Traditional Recruitment maneuvers	⚠ CONSIDER: Prone ventilation 12-16 h
⚠ CONSIDER: Conservative fluid strategy	⚠ CONSIDER: Prone ventilation 12-16 h	⚠ CONSIDER: STOP if no quick response A trial of inhaled Nitric Oxide
⚠ CONSIDER: Empiric antibiotics	⚠ CONSIDER: if proning, high P_{plat} , asynchrony NMBA infusion for 24 h	⚠ CONSIDER: follow local criteria for ECMO V-V ECMO or referral to ECMO center
? Uncertain: Systematic corticosteroids	🚫 Don't do: Staircase Recruitment maneuvers	
	⚠ CONSIDER: Short course of systemic corticosteroids	
	? Uncertain: Antivirals, chloroquine, anti-IL6	