



Management of Acute Hypercapnic Respiratory Failure: Case Discussion



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**Which one is the
patient with
hypercapnia?**



ACUTE HYPERCAPNIC RESPIRATORY FAILURE (AHRF)

- ✓ The inability to provide alveolar ventilation to keep PaCO₂ levels within normal limits
- ✓ pH <7.35 and PCO₂ > 45 mmHg indicate acute respiratory acidosis
- ✓ Seen in 20% of COPD exacerbations
- ✓ 12% of hypercapnic COPD cases are lost at the first admission

*Roberts CM, Stone RA, Buckingham RJ, et al Thorax
2011;66:43*

ACUTE HYPERCAPNIC RESPIRATORY FAILURE (AHRF)

- ✓ Mortality is high in hypercapnic asthma attacks
- ✓ Poor prognostic indicator in complicated cystic fibrosis and bronchiectasis cases
- ✓ It can occur suddenly in neuromuscular and chest wall related diseases, acute decompensated attacks are common in chronic conditions and is a precursor for long-term NIMV at home
- ✓ If not recognized or treated, it can cause seizures, coma, arrest and death

Hypercapnia

- Hypercapnia is always an indicator of inadequate ventilation.
- PaCO₂;
 - Directly proportional to CO₂ production
 - Inversely proportional to alveolar ventilation

$$\dot{V}_E = f \times V_T$$

$$\dot{V}_E = \dot{V}_A + \dot{V}_D$$

$$\dot{V}_A = \dot{V}_E - \dot{V}_D$$

$$PACO_2 = \frac{\dot{V}CO_2 \times 0.863}{\dot{V}_A}$$

$$pCO_2 = k \times \frac{V_{CO_2}}{V_E (1 - V_D/V_T)}$$

Causes of Respiratory Failure

Failure to Ventilate

Neurological

Respiratory Center

Opioids, Anesthetics, Brain Injuries

Cervical Nerves C3,4,5

Spinal Injuries

Phrenic Nerves

Chest trauma, Surgery

Neuromuscular Junction

Neuromuscular Blockers

Myasthenia Gravis

Muscular

Myopathy

Steroids

Myasthenia Gravis

Polyneuropathy/Polymyopathy
of Critical Illness

Diaphragm
Intercostals

Failure to Maintain Airway

Failure of Gas Flow:

Airway Obstruction

-Upper: teeth, tongue

-Glottic:

laryngeal edema

laryngospasm

-Lower: bronchospasm

Inhaled objects

Chest Wall

Flail Chest

Pleural Cavity

Pneumothorax

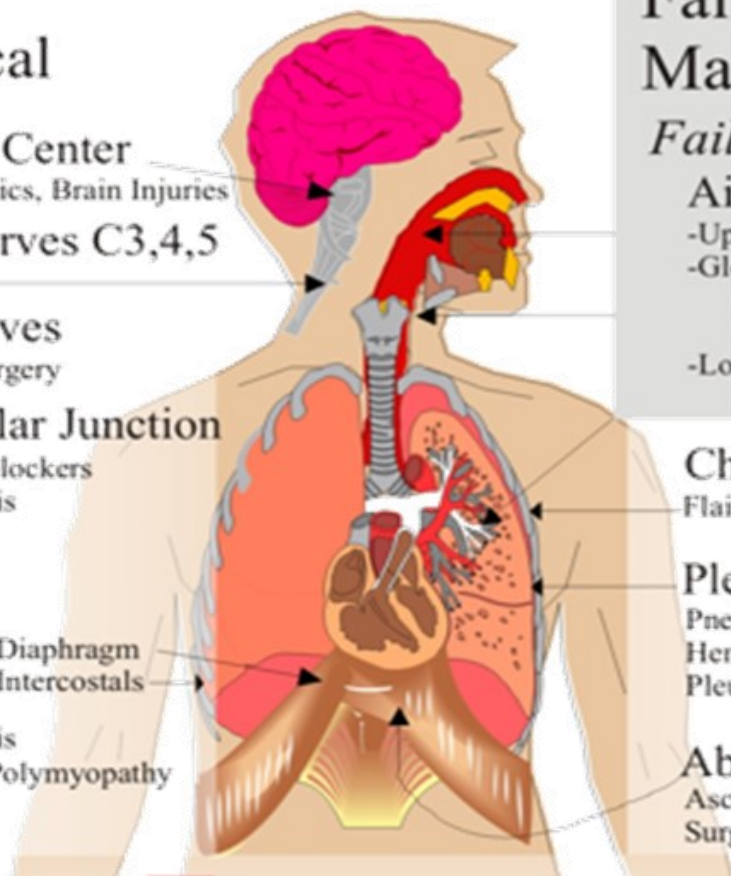
Hemothorax

Pleural Effusion

Abdominal Compression

Ascites/Hemoperitoneum

Surgical Packs etc



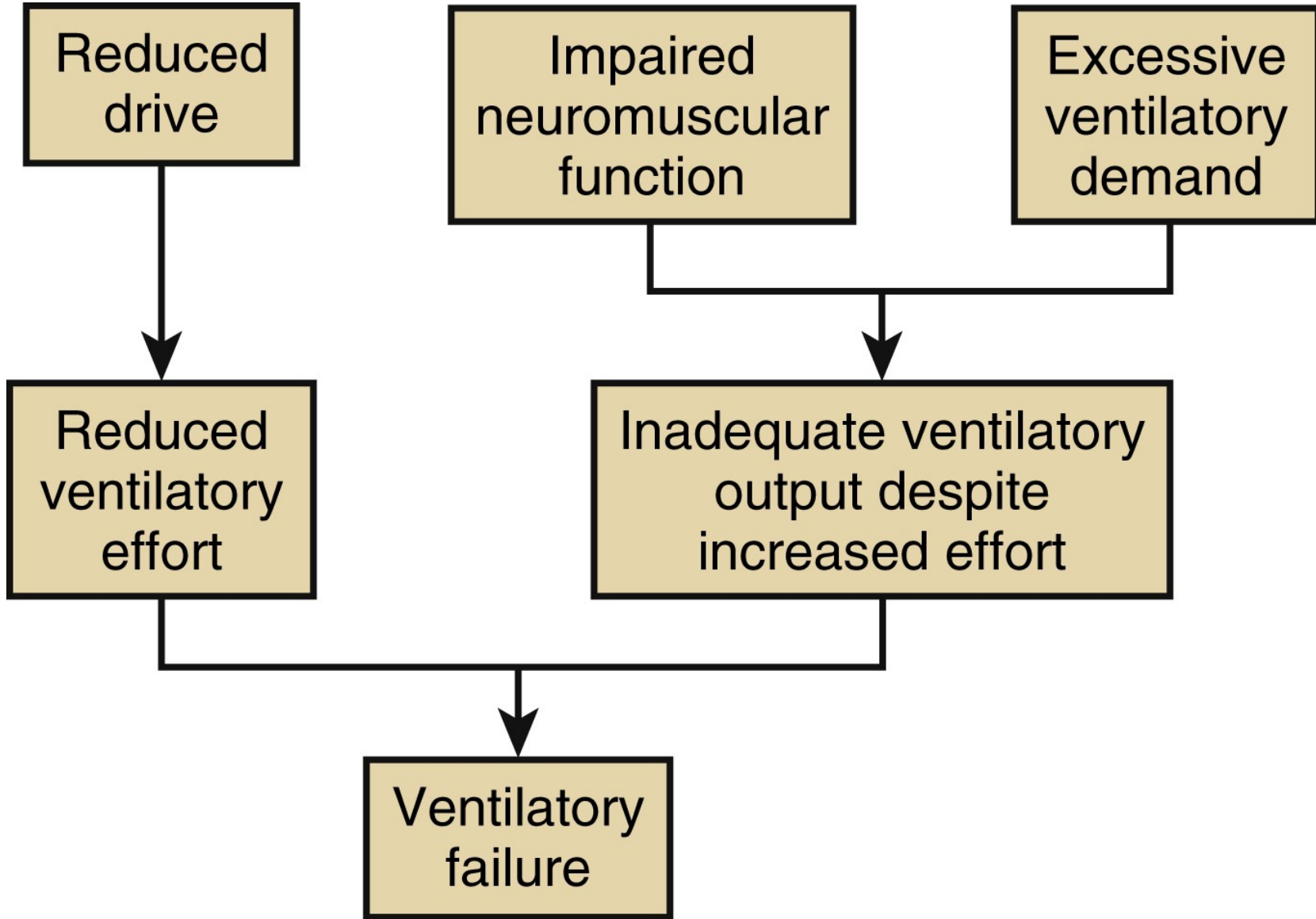


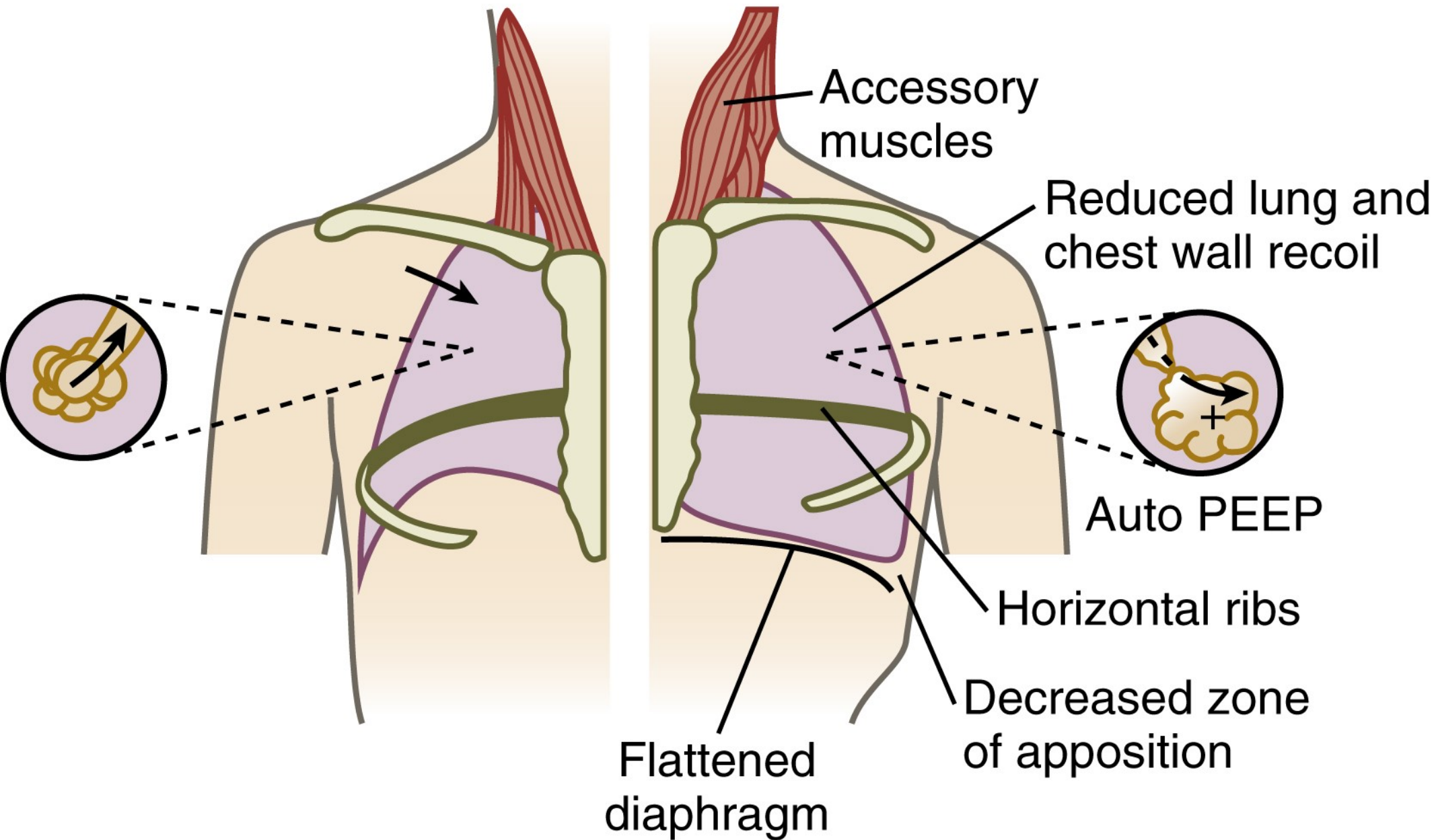
Table 132.1 Clinical Classification of Ventilatory Failure by Site

Site of Defect	Mechanism or Type	Clinical Examples
VENTILATORY DRIVE	Congenital	Primary alveolar ventilation (Ondine's curse)
	Acquired	Drug overdose (opioids, sedatives, alcohol); propofol; cerebrovascular accident; neoplasm; carotid body resection
	Combination	Obesity-hypoventilation syndrome; myxedema
NEURAL TRANSMISSION		
Spinal cord	Trauma	Cervical spinal cord injury
	Vascular	Vascular accident
	Tumor	Primary or metastatic
	Other	Poliomyelitis; amyotrophic lateral sclerosis
	Demyelinating	Acute idiopathic demyelinating polyneuropathy (Guillain-Barré syndrome)
Peripheral nerves	Phrenic nerve lesion	Trauma; cardiac surgery; neoplasm; idiopathic
Neuromuscular junction	Autoimmune	Myasthenia gravis
	Infectious/toxins	Botulism, tetanus, tick paralysis
	Pharmacologic	Neuromuscular blocking agents
	Critical illness	ICU-acquired weakness, ventilator-induced diaphragmatic dysfunction

VENTILATORY MUSCLES	Congenital Autoimmune Acquired	Muscular dystrophy Polymyositis; dermatomyositis Hypophosphatemia; hypokalemia; hypomagnesemia; myxedema
THORACIC		
Vertebrae and rib cage	Decreased mobility	Kyphoscoliosis; tight casts or bandages; ankylosing spondylitis; flail chest
Soft tissues	Extrapulmonary restriction and decreased mobility	Severe obesity
Pleura	Extrapulmonary restriction	Pneumothorax; pleural effusion; pleural thickening; malignancy
AIRWAYS		
Upper	Obstruction	Epiglottitis, foreign body, tumor, vocal cord paralysis, tracheomalacia
Lower	Obstruction	COPD, acute severe asthma
PARENCHYMA	Increased dead space and very high \dot{V}/\dot{Q} Very low \dot{V}/\dot{Q} ; shunt	COPD Severe ARDS
PULMONARY CIRCULATION	General hypoperfusion Localized hypoperfusion	Hypovolemic or cardiogenic shock, CPR, pulmonary hyperinflation (intrinsic PEEP) Pulmonary thromboembolism; venous air embolism
OTHER	Increased CO ₂ production (inflammation; hypermetabolism; muscle activity) Exogenous CO ₂ inhalation	Fever; sepsis; burns; severe trauma; shivering; tetany; seizures; malignant hyperthermia Laboratory or industrial accident; therapeutic use; rebreathing

Normal

COPD



ARTERIAL BLOOD GASES ANALYSIS

- ✓ In acute respiratory acidosis, PaCO_2 is >45 mmHg while pH is <7.35
- ✓ In acute exacerbation on a chronic basis, pH is higher than expected
- ✓ In chronic hypercapnia, pH is normal or close to normal

- ✓ To distinguish between global hypoventilation and pulmonary disease «**A-a gradient**» should be calculated ($\text{PAO}_2 - \text{PaO}_2$)
- ✓ It can be calculated practically with the formula **Age x 0.3**
- ✓ If PCO_2 is high and **A-a gradient is normal**, it means there is **hypoventilation**
- ✓ If PCO_2 is high and **A-a gradient >20 mmHg**, it indicates underlying **lung disease**

MECHANICAL VENTILATION IN HYPERCAPNIC RESPIRATORY FAILURE

WHAT IS THE PURPOSE?

- ✓ Reducing inspiratory muscle and respiratory load
- ✓ Reducing dynamic hyperinflation and auto-PEEP
- ✓ Improving gas exchange

Non-invasive mechanical ventilation (NIMV)

Invasive mechanical ventilation (IMV)

Hemodynamic instability
Loss of consciousness
Inability to protect airway, inability to expel secretions
History of facial, esophageal, gastric surgery

How to manage the patient?



NIV



HFNC



IMV

When should high flow nasal cannula (HFNC) be used in the clinical setting?

Hypoxemic respiratory failure

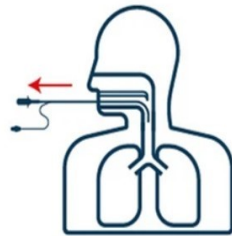
(moderate certainty)



**Strong
recommendation**

Following extubation

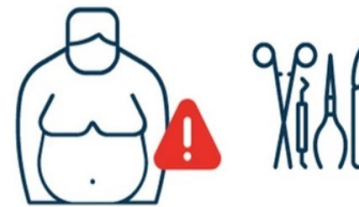
(moderate certainty)



**Conditional
recommendation**

**Postoperative HFNC in high risk
and/or obese patients following
cardiac or thoracic surgery**

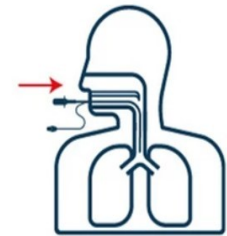
(moderate certainty)



**Conditional
recommendation**

Peri-intubation period

(moderate certainty)



**No
recommendation**



The role for high flow nasal cannula as a respiratory support strategy in adults: a clinical practice guideline. *Intensive Care Med* 46, 2226–2237 (2020).

In hypercapnic ventilatory failure;

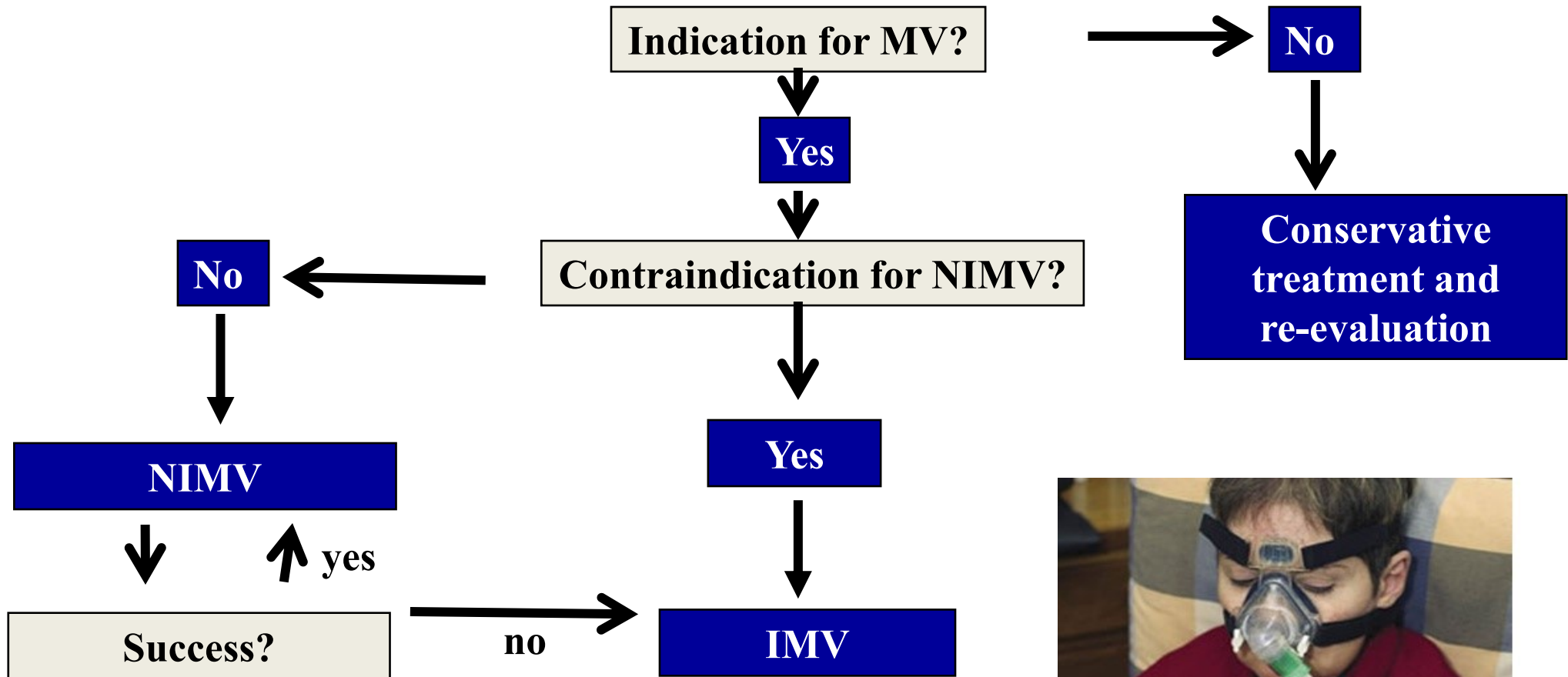
- ✓ First choice for hypercapnic respiratory failure NIMV
- ✓ HFNC may be an alternative for patients with low tolerance and mild to moderate respiratory acidosis
- ✓ HFNC alternately during NIMV treatment

Use of nasal high flow oxygen during acute respiratory failure. Intensive Care Med 46, 2238–2247 (2020).

High-Flow Nasal Cannula for Chronic Obstructive Pulmonary Disease with Acute Compensated Hypercapnic Respiratory Failure: A Randomized, Controlled Trial. Int J Chron Obstruct Pulmon Dis. 2020 Nov 24;15:3051-3061.

Huang Y, Lei W, Zhang W, Huang JA. High-Flow Nasal Cannula in Hypercapnic Respiratory Failure: A Systematic Review and Meta-Analysis. Can Respir J. 2020;2020:7406457. Published 2020 Oct 29. doi:10.1155/2020/7406457

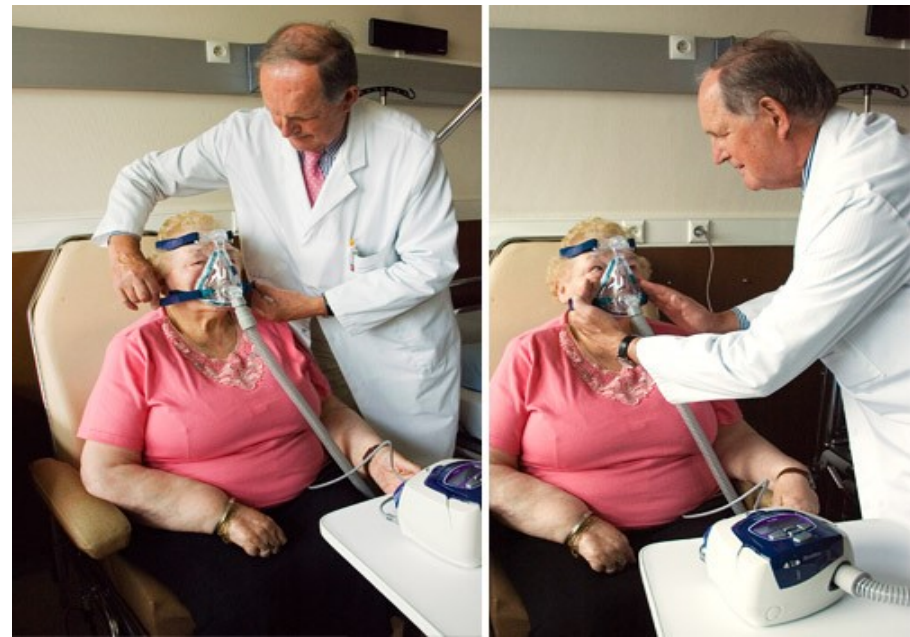
Mechanical Ventilation



INDICATIONS FOR NIMV

- ✓ Moderate to advanced dyspnea
- ✓ Tachypnea
- ✓ Accessory respiratory muscle use
- ✓ Gas exchange impairment:

$\text{PaCO}_2 > 45 \text{ mmHg}$, $\text{Ph} < 7.35$



!!! Hypercapnic encephalopathy is not a contraindication for NIMV !!!

CONTRAINDICATIONS FOR NIMV:

ABSOLUTE:

- Respiratory arrest
- Difficulty fitting mask



PARTIAL:

- Medically unstable conditions
- Hypotensive shock
- Uncontrolled
- Cardiac ischemia
- Arrhythmia
- Upper GI bleeding
- Agitation, inability to communicate
- Inability to protect airway
- Swallowing disorder
- Hypersecretion
- Multiple organ failure
- Recent upper GI or GI surgery



97 HR bpm 117/79 BP mmHg 89 SpO2 % 26 RR BrPm

100% Prog1 ST [Lock Icon]

Clinical Settings 1/3

Pathology Normal	Mode ST	IPAP 10.0 cmH2O
EPAP 4.0 cmH2O	Backup Rate 10 bpm	Advanced Settings

Lk 0 | RR 26 | Ti 0.6 | Vt 239 | MV 6.3

[Lung Icon] [RR Icon] [Info Icon] [Power Icon]

Bedside


ABG



Suggested appropriate initial ventilator settings for a COPD patient

Click on the settings button to see the reason why this setting is chosen



Setting	Appropriate value in COPD	Why
Mode	S/T mode	 <p>Setting cycle sensitivity to HIGH results in shorter inspiratory time and longer expiratory time useful to ensure complete lung emptying and avoid hyperinflation. MEDIUM trigger sensitivity is sufficient in obstructive lung disease.</p>
IPAP	12-14 cm	
EPAP or PEEP	4-6 cm	
Back up rate	12 bpm	
Ti Min Ti Max	0.3 sec. 1.0 sec	
Trigger Cycle	Medium High	
Rise Time Fall Time	150 ms 200 ms	

Initial settings in acute neuromuscular disease

The Amyotrophic Lateral Sclerosis is a restrictive lung disease.
The suggested initial settings are as shown:



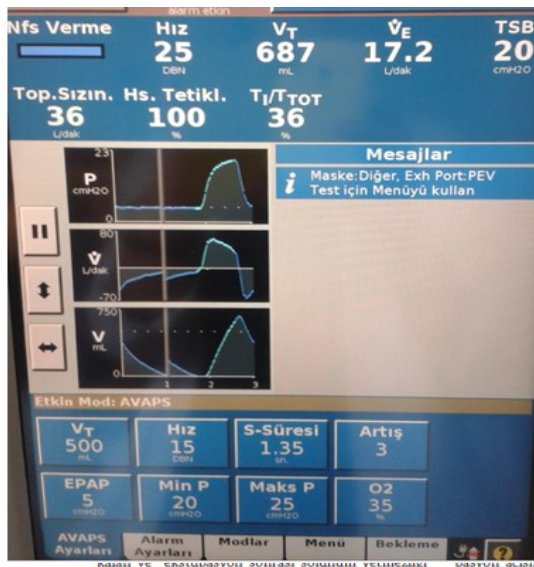
Mode	S/T
IPAP	11 cm H ₂ O
EPAP	5 cm H ₂ O
Backup rate	12 bpm
Ti Min	0.5 sec
Ti Max	1.5 sec
Trigger	Medium
Cycle	Low
Rise time	200-300 msec
Fall time	200 msec
Oxygen	0 - 4 LPM

AVAPS

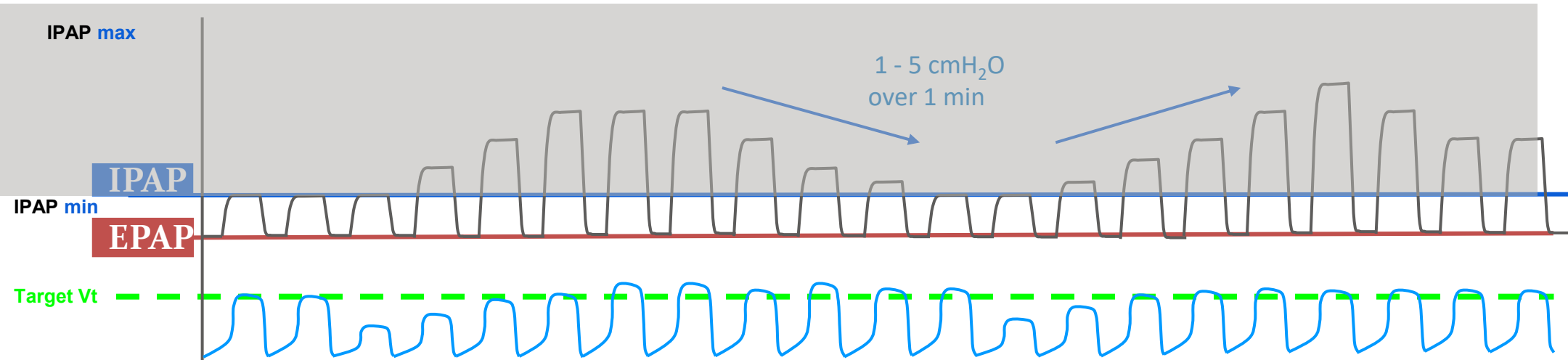
Average Volume Assured Pressure Support Volume Guaranteed Pressure Support

1. Measuring V_{te} in each breath
2. Comparing the average V_{te} with the target V_{te} in each breath
3. Calculating the pressure required to reach the target V_{te}
4. To do this, progressively increasing the IPAP value in each breath if necessary
($< 1\text{cmH}_2\text{O}/\text{min}$)

It is a system that automatically adjusts the pressure to ensure that the patient has an average good ventilation.



AVAPS



IVAPS

Intelligent Volume Assured Pressure Support

Target alveolar ventilation

$V_a : f \times (V_T - V_d) : 12 \times (500 - 150) \text{ ml} : 4,2 \text{ lt/dk}$

Target V_a , EPAP, Height, f , Min PS, Max PS

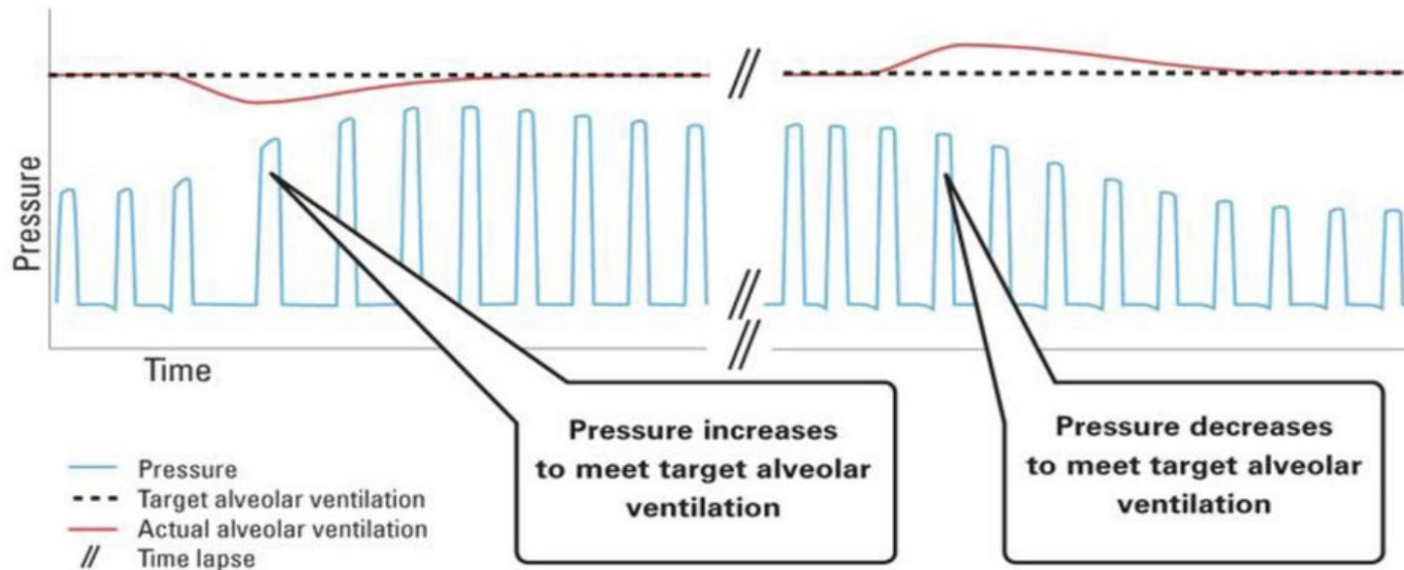
IVAPS

Intelligent Volume Assured Pressure Support

Fixed alveolar ventilation

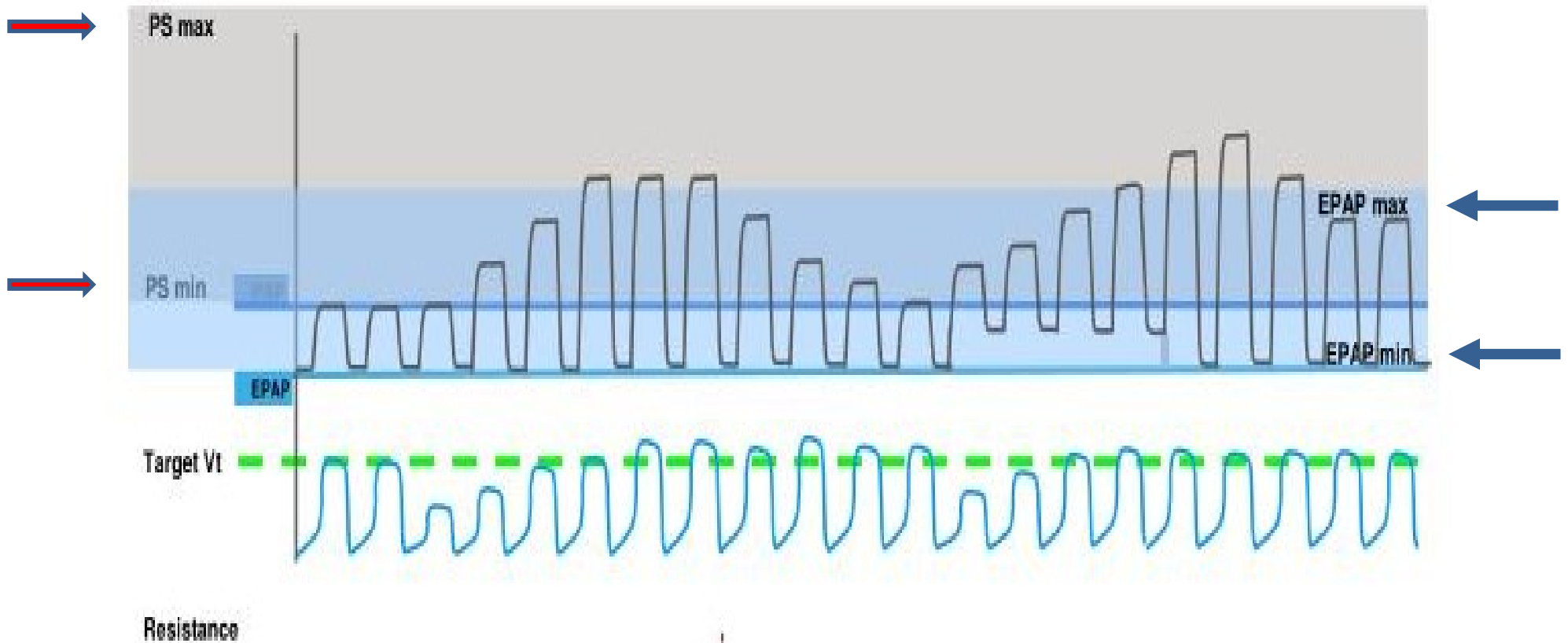
Automatically adjusts pressure support and respiratory rate to provide optimal ventilatory support

Intelligently changing pressure support to maintain alveolar ventilation



AVAPS-AE

Noninvasive auto-titrating ventilation



AVAPS-AE

For Whom?

- ✓ Neuromuscular diseases
- ✓ Restrictive thoracic wall pathologies
- ✓ COPD patients with chronic respiratory failure
- ✓ Chronic hypercapnic obesity hypoventilation patients

Indications for NIV

COPD

pH < 7.35
pCO₂ > 6.5
RR > 23

If persisting after bronchodilators and controlled oxygen therapy

Neuromuscular disease

Respiratory illness with RR > 20 if usual VC < 1L even if pCO₂ < 6.5
Or
pH < 7.35 and pCO₂ > 6.5

Obesity

pH < 7.35, pCO₂ > 6.5, RR > 23
Or
Daytime pCO₂ > 6.0 and somnolent

Contraindications for NIV

Absolute

Severe facial deformity
Facial burns
Fixed upper airway obstruction

Relative

pH < 7.15
(pH < 7.25 and additional adverse feature)
GCS < 8
Confusion/agitation
Cognitive impairment (warrants enhanced observation)

Indications for referral to ICU

AHRF with impending respiratory arrest

NIV failing to augment chest wall movement or reduce pCO₂

Inability to maintain SaO₂ > 85-88% on NIV

Need for IV sedation or adverse features indicating need for closer monitoring and/or possible difficult intubation as in OHS, DMD.

NIV SETUP

Mask

Full face mask (or own if home user of NIV)

Initial Pressure settings

EPAP: 3 (or higher if OSA known/expected)

IPAP in COPD/OHS/KS 15 (20 if pH < 7.25)

Up titrate IPAP over 10-30 mins to IPAP 20-30 to achieve adequate augmentation of chest/abdo movement and slow RR

IPAP should not exceed 30 or EPAP 8* without expert review

IPAP in NM 10 (or 5 above usual setting)

Backup rate

Backup Rate of 16-20. Set appropriate inspiratory time

I:E ratio

COPD 1:2 to 1:3
OHS, NM & CWD 1:1

Inspiratory time

0.8-1.2s COPD
1.2-1.5s OHS, NM & CWD

Use NIV for as much time as possible in 1st 24 hours.
Taper depending on tolerance & ABGs over next 48-72 hours

SEEK AND TREAT REVERSIBLE CAUSES OF AHRF

NIV Monitoring

Oxygenation

Aim 88-92% in all patients

Note: Home style ventilators CANNOT provide > 50% inspired oxygen.

If high oxygen need or rapid desaturation on disconnection from NIV consider IMV.

Red flags

pH < 7.25 on optimal NIV
RR persisting > 25

New onset confusion or patient distress

Actions

Check synchronisation, mask fit, exhalation port: give physiotherapy/bronchodilators, consider anxiolytic

CONSIDER IMV

NIV Not indicated

Asthma/Pneumonia

Refer to ICU for consideration IMV if increasing respiratory rate/distress
or
pH < 7.35 and pCO₂ > 6.5

* Possible need for EPAP > 8

Severe OHS (BMI > 35), lung recruitment eg hypoxia in severe kyphoscoliosis, oppose intrinsic PEEP in severe airflow obstruction or to maintain adequate PS when high EPAP required

Ventilator Settings

- Beginning:
 - EPAP: 3-5 cmH₂O
 - IPAP: 8-12 cmH₂O
 - PS: 7-16 cmH₂O
- Change:
 - EPAP + 1 (according to SpO₂)
 - IPAP + 2 (according to TV and PaCO₂)
- Patient comfort:
 - Rise time: 0.1 sn
 - Inspiryum time: <1.0 sn.
- Oxigenation: Target SpO₂<90-92

Patient 1

56 year old female, Height: 160, Weight: 85

Symptoms: Respiratory distress, cough increasing in the last 5 days

Has Severe COPD

Has not used nebulized bronchodilators in the last 3 days

Chest X-ray: Increased ventilation, infiltrations compatible with pneumonia

Blood gases analyses: pH: 7.26, pCO₂: 82, pO₂: 75, HCO₃: 20, SO₂: 90

CRP: 20 mg/L BK: 17,000 Creatine: 1.8 mg/dl BUN: 55

Physical examination:

**Uses accessory
respiratory muscles**

**Respiratory rate:
27/min**

**Auscultation:
Quiet lung**

**Vital findings:
Temperature: 38.5
Pulse: 125/min
BP: 150/80**



COPD Exacerbation with respiratory failure



NON-INVASIVE MECHANICAL VENTILATION



When should we perform the first blood gas check?
Should we transfer the patient to intensive care?

- ✓ **The patient should be closely monitored for the first eight hours**
- ✓ **It is expected that pH and PaCO₂ will improve within 30 minutes-2 hours**
- ✓ **NIV can also be applied in a short-term ward environment**

Patient 1

Admission ABG: pH: **7.26**, pCO₂: **82**, pO₂: 75, HCO₃: 20, SO₂: 90



IPAP: 12, EPAP:6 , fiO₂:4 lt/dk, rise time:0.3 sn, exp trigger:%45

After 1 hour: pH: **7.28**, pCO₂: **78**, pO₂: 80, HCO₃: 21, SO₂: 93



IPAP: 16, EPAP:7 , fiO₂:4 lt/dk, rise time:0.1 sn, exp trigger:%75

NIV is continued

How to set the NIV?

- ✓ **IPAP** →
- ✓ **EPAP (CPAP/PEEP)** →
- ✓ **Rise time** →
- ✓ **Time for inspiration** →
- ✓ **Trigger** →
- ✓ **Expiratory trigger** →
- ✓ **Respiratory rate for S/T mode** →

- ✓ **AVAPS – TV, Min IPAP, Max IPAP**
- ✓ **Alarm settings!!**

IPAP= (PS+PEEP) or (PS+EPAP)

6-8 mL/kg TV

Preventing CO2 rebreathing,
minimum 4-5 cmH2O

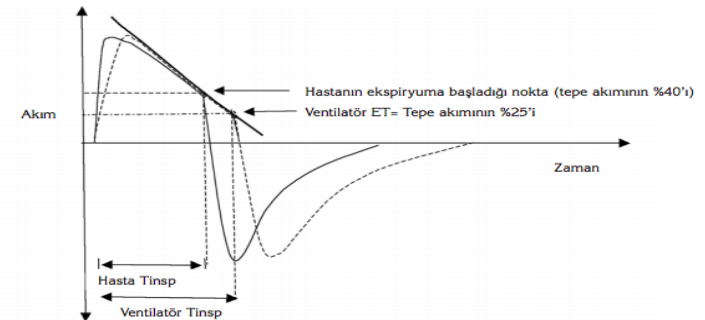
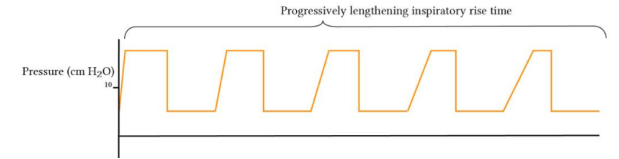
Short 0.1 sec

0.8-1.0 sec

According to patient effort and comfort

For the COPD patients %50-75

12-16/min



GENERALLY NIMV START-UP SETTINGS



The purpose of PEEP during NIMV is to overcome intrinsic PEEP in obstructive diseases

Since PEEP cannot be measured during NIMV, low PEEP (≤ 5 cmH₂O) is recommended



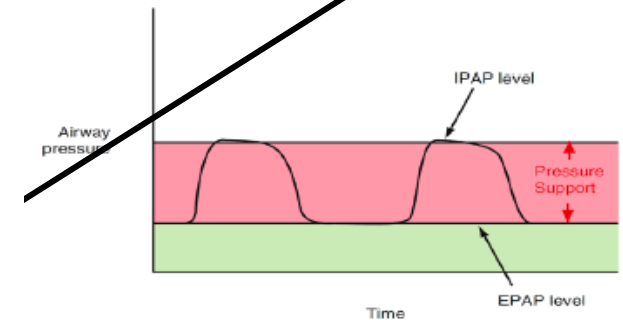
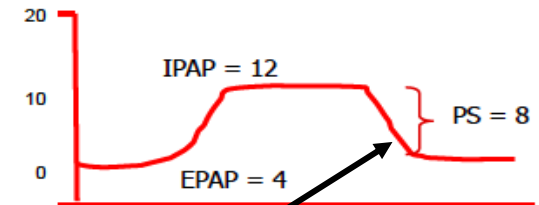
Ventilatör Basınç Ayarları



IPAP:
8-10 cm H₂O
Minute ventilation

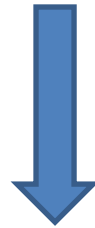
EPAP:
4-5 cm H₂O
Oxygenation

Difference between PS
7-16 cm H₂O



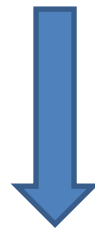
Patient 1

After 2 hour of NIMV: pH: 7.27, pCO₂: 84, pO₂: 70, HCO₃: 19, SO₂: 85



IPAP: 22, EPAP:8 , fiO₂:6 lt/dk

After 30 min : The patient uses accessory respiratory muscles, SS: 30, agitated, sweating, confused consciousness



Intubation and Invasive mechanical ventilation

UNTIL WHEN SHOULD WE APPLY NIMV?

BTS/ICS guideline for the ventilatory management of acute hypercapnic respiratory failure in adults

- ✓ Until the patient's general condition improves and **pH and pCO₂** return to normal
- ✓ NIV duration should be **maximized in the first 24 hours** as tolerated by the patient
- ✓ Depending on the patient's own respiratory effort and pCO₂ level, its frequency is reduced within 2-3 days before stopping at night

Patient 1

Patient is being monitored with IMV for 4 days under maximum medical treatment



4th day blood gas: pH: 7.38, pCO₂: 55, pO₂: 90, HCO₃: 29, SO₂: 96



SBT failed, but patient is cooperative, Chest X-ray regressed



Should we extubate? Should we change MV settings?

Official ERS/ATS clinical practice guidelines: noninvasive ventilation for acute respiratory failure

Recommendations

We suggest NIV be used to facilitate weaning from mechanical ventilation in patients with hypercapnic respiratory failure. (Conditional recommendation, moderate certainty of evidence.)

We do not make any recommendation for hypoxaemic patients.

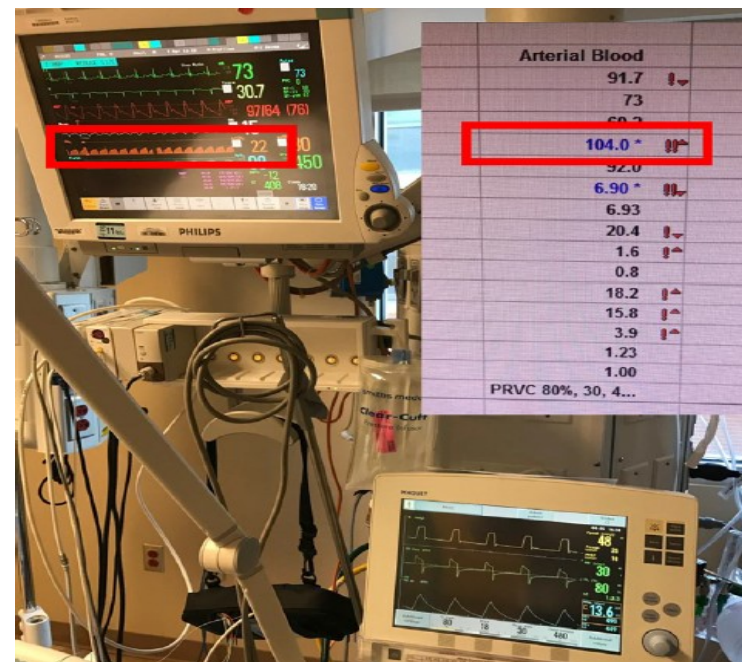
- **NIV after extubation in patients with acute exacerbations of COPD who fail SBT****
- **The use of NIV for weaning is complex and requires more experience**
- **Therefore, the committee offers a conditional recommendation rather than a strong recommendation**

Patient 1

- The patient was extubated on the 4th day of his ICU stay and after continuing his treatment with NIV for a while, she was discharged in stable condition.

HOW SHOULD MV SETTINGS BE?

- For NIV
 - Increase the gradient between IPAP and EPAP
- For IMV
 - Initially 100% FiO₂
 - PEEP 5, TV 6 to 8 mL / kg PBW
 - If there is spontaneous breathing, expirium trigger should be adjusted
 - Respiratory rate 14-16/min
 - Auto-PEEP monitoring should be done, measured with expirium hold
 - If auto PEEP is not available, respiratory rate is gradually increased



Patient 2

- HD, Age: 60, gardener,
- Brought to the emergency room due to difficulty breathing, weakness, and confusion.
- Has been feeling weak for a year, complaints have increased in the last 3 weeks, cannot cough strongly
- Uses bronchodilators due to COPD
- Diagnosed with Diabetes
- Physical examination does not have much obstruction, but sounds are very reduced. Heart rhythmic, no edema, no cyanosis.

Patient 2

Admission to ER

PaO₂ :58

PaCO₂: 68

pH : 7.28

O₂ Sat : 87

HCO₃ : 26

BE: 3





Patient 2

- $D(A-a)O_2 = 8$ mmHg
- Questioned for intoxication.
- Thyroid function tests are normal
- Nutrition is good
- No electrolyte imbalance
- Left ventricular functions are normal
- OSA is not considered

Patient 2

	<u>Admission</u>	<u>Bronchodilator+</u> <u>Oxygen</u>	<u>NIV 1h</u>	<u>NIV 2h</u>
PaO ₂	: 58	: 61	: 82	: 79
PaCO ₂	: 68	: 71	: 59	: 50
pH	: 7.28	: 7.31	: 7.38	: 7.41
O ₂ Sat	: 87	: 89	: 96	: 96
HCO ₃	: 26	: 32	: 35	: 32
BE	: 3	: 11	: 9	: 8

Patient 2

- Neurology consultation
- Cerebral MRI:
 - lacunar infarct
- Tensilon and Pridostigmine tests are normal
- EMG:
 - Compatible with Motor Neuron Disease

Patient 2

- Spirometry:
 - FVC: 1.98 lt (%67)
 - FEV₁: 1.59 lt (%67)
 - FEV₁/FVC: 80
- BiPAP
- EPAP 5, IPAP 12

Patient 2

- Motor neuron disease
- Spinal cord anterior horn disease
- **ALS:** Amyotrophic Lateral Sclerosis
- Starts from the distal ends of the upper extremities
- Weakness, atrophy, fasciculations, inability to perform fine movements, dysphagia, dysarthria, Babinski positivity,
- The most important feature is the absence of sensory deficit.
- Respiratory muscle involvement
- Inability to cough, difficulty swallowing, basal atelectasis, restrictive loss

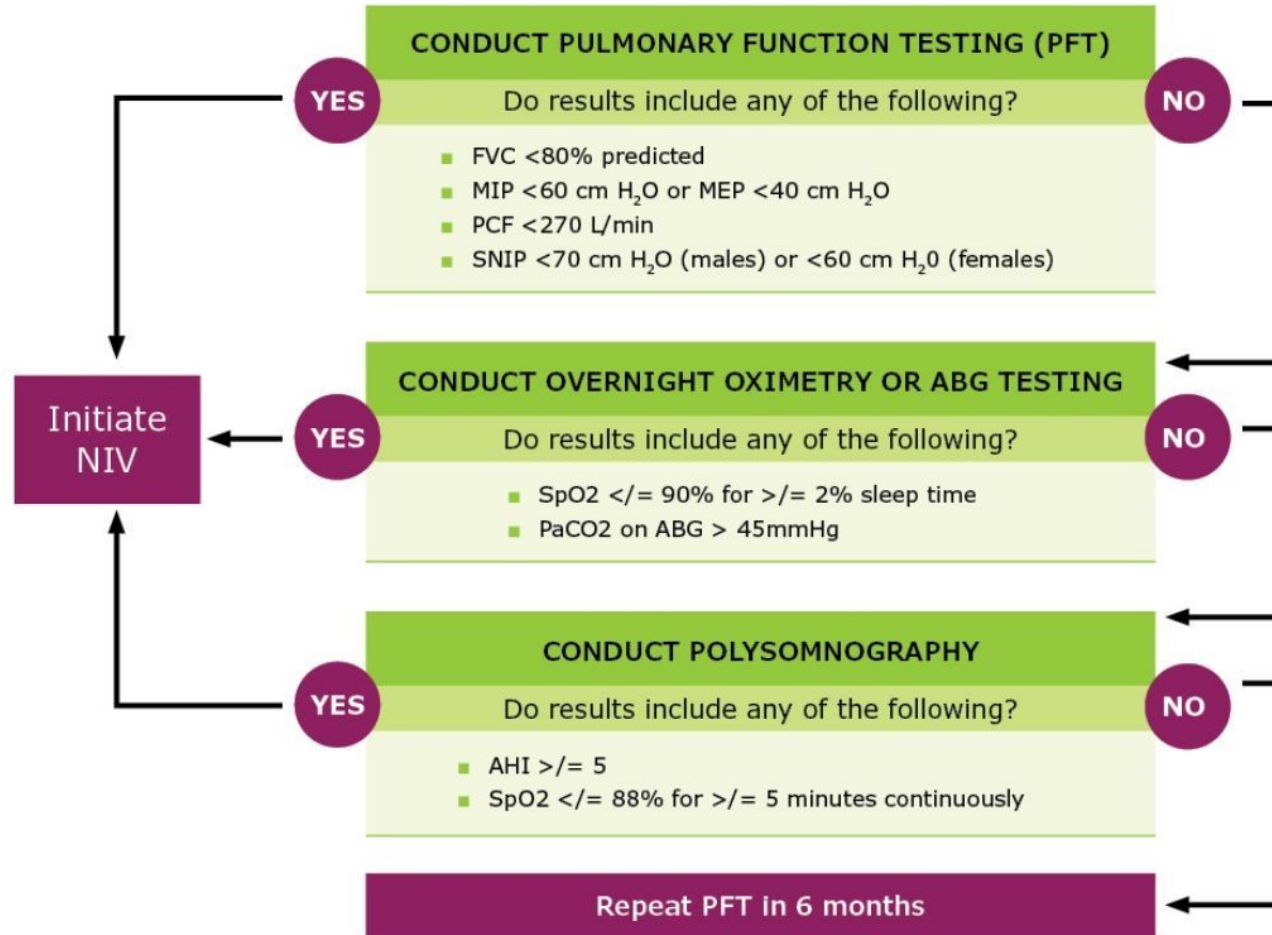
Respiratory Management of Patients With Neuromuscular Weakness



An American College of Chest Physicians Clinical Practice
Guideline and Expert Panel Report

Algorithm for Initiation of Noninvasive Ventilation

Based on a CHEST clinical practice guideline and expert panel report, this flowchart outlines an algorithm for initiation of noninvasive ventilation (NIV) for adult patients with neuromuscular disease experiencing fatigue, headache, concentration/memory difficulty, and/or respiratory symptoms.



Conclusion

- Hypercapnic respiratory failure treatment is arranged according to the underlying cause
- NIV is the most successful treatment method
- Early weaning should be tried to avoid IMV complications
- Long-term home treatments should be arranged for patients with chronic hypercapnia and symptoms