

1 **PATIENT-VENTILATOR ASYNCHRONY IN A TRAUMATICALLY INJURED**
2 **POPULATION**

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1 **Short Title: Asynchrony in Trauma Patients**

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1 ABSTRACT

2 **Introduction:** Prolonged mechanical ventilation, increased length of hospital stay, and a lower
3 rate of home discharge have been reported with patient-ventilator asynchrony in medical
4 patients. Though commonly encountered, asynchrony is poorly defined within a traumatically
5 injured population.

6 **Methods:** Mechanically ventilated trauma patients at an urban, level I center were enrolled.
7 Breath waveforms were recorded over 30 minutes within the first 48 hours following intubation.
8 Asynchronous breaths were defined as ineffective patient triggering, double triggering, short
9 cycle breaths, and long cycle breaths. Asynchronous patients were defined as having asynchrony
10 in $\geq 10\%$ of total breaths. Demographic, injury, sedation/delirium scores, clinical and discharge
11 outcomes were prospectively collected.

12 **Results:** 35 patients were enrolled. Median age was 47 years, 77.1% male, 28.6% with
13 penetrating injuries, 16% with a history of COPD, median ISS of 22 (IQR 17-27), and a median
14 chest AIS of 2 (IQR 0-6). 15,445 breaths were analyzed. Asynchrony was present in 25.7% of
15 patients. No statistical differences between asynchronous and non-asynchronous patients were
16 found for age, sex, injury mechanism, COPD history, delirium/sedation scores, PaO₂/FiO₂
17 ratios, PEEP, blood gas values, sedative, narcotic and haloperidol use. Asynchronous patients
18 more commonly used synchronized intermittent mandatory ventilation (SIMV) (100% vs.
19 38.5%; p=0.002) and took lower median spontaneous breaths per minute (4 (IQR 3-8) vs. 12
20 (IQR 9-14); p=0.007). SIMV with set respiratory rates ≥ 10 breaths per minute were associated
21 with increased asynchrony rates (85.7% vs. 25.0%, p=0.02). We found no difference in
22 ventilator days, ICU and hospital lengths of stay, % discharged home, or mortality between
23 asynchronous and non-asynchronous patients.

24 **Conclusions:** Ventilator asynchrony is common in trauma patients. It may be associated with
25 SIMV with a set respiratory rate ≥ 10 breaths per minute, though not with increased duration of
26 mechanical ventilation, length of stay, or discharge disposition.

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28 **Key Words:** Mechanical ventilation, asynchrony, trauma, critical care

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33 **INTRODUCTION**

1 Patient-ventilator asynchrony is a common occurrence in critically ill intensive care unit
2 (ICU) patients. Asynchrony can be defined as a mismatch between the patient's ventilator needs
3 and the ventilator settings and operating characteristics (1-5). Studies have shown that
4 asynchrony may be associated with a longer duration of mechanical ventilation, due to
5 ineffective weaning, resulting in increased ICU and hospital length of stay and poor outcomes
6 (1,6).

7 Asynchrony has been evaluated in a number of studies utilizing review of the pressure,
8 volume and flow waveforms from the mechanical ventilator (1-5). This technique has
9 limitations which include the ability to visualize the waveforms on a variety of different
10 ventilators with different display resolutions, reliance on the internal measurement systems of the
11 ventilator, and lack of traditional esophageal pressure monitoring. Nonetheless, previous work
12 has demonstrated the validity of ventilator waveforms evaluation compared to esophageal
13 pressure monitoring for the detection of missed triggers and asynchrony (1). Additionally, these
14 waveforms are commonly utilized by the clinical staff to both detect and alleviate asynchrony at
15 the bedside of patients experiencing respiratory distress.

16 Traumatically injured patients differ greatly from previously investigated populations by
17 virtue of lower ages with resultant decreased comorbidities; nonetheless, bedside clinicians
18 encounter asynchrony. To date, there have been no prospective studies of asynchrony in trauma
19 patients thus the application of previous conclusions made in medical patients appear to be
20 speculative. We conducted a prospective study to determine the frequency and characteristics of
21 asynchrony in a traumatically injured patient population. We hypothesize that asynchrony is
22 present in trauma patients though the characteristics and implications upon outcomes are
23 different than the published data in populations of medically ill patients.

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2 **METHODS**

3 **Patient Population**

4 All traumatically injured patients in the surgical intensive care unit (ICU) of the
5 University Hospital that had spontaneous respiratory efforts within 48-hours of initiation of
6 mechanical ventilation were eligible for study inclusion. The University Hospital of the
7 University of Cincinnati is an adult American College of Surgeons-verified Level I trauma center
8 that serves 1.8 million people in southwestern Ohio, northern Kentucky and southeastern
9 Indiana. The hospital is a 693-bed facility with over 100 critical care beds, 34 of which are
10 dedicated to injured and critically ill surgical patients. Exclusion criteria for study participation
11 were age < 18 years old, patients without spontaneous respirations due to injury, sedation, or
12 chemical paralysis, patients with air leaks in the ventilator circuit such to preclude accurate data
13 collection of pressure-time, flow-time and tidal volume-time waveforms, and use of a ventilator
14 other than the Draeger Evita XL (Draeger Medical, Telford, PA). The presence or absence of
15 asynchrony was not a criterion for study inclusion. Ventilator-patient waveforms were not
16 collected before study participation. The Institutional Review Board of the University of
17 Cincinnati approved the study with informed consent obtained from the patients legally
18 authorized representative. This study was registered with ClinicalTrials.gov with an identifier of
19 NCT01049958.

20 **Data Acquisition**

21 Real time pressure, flow, and volume waveforms generated by the patient were acquired
22 and recorded over 30 minutes by a laptop computer connected to the RS 232 output port of the

1 ventilator for those patients that met inclusion criteria. Recordings were done once per patient
2 during the first 48-hours of ventilator initiation with none of the patients having a tracheostomy.
3 Ventilator settings were determined by the clinicians caring for the patient and were not
4 manipulated during the recording period. Respiratory therapists provided usual care during the
5 period of recording with all patients having a heat moisture exchanger (HME). Recordings were
6 performed by study personal during periods in which healthcare delivery did not preclude
7 investigation. Waveforms were analyzed at a later date utilizing Vent View software (Vent
8 View, Draeger Medical, Telford, PA).

9 **Asynchrony Criteria**

10 Asynchronous breaths included ineffective triggered breaths, double-triggered breaths, as
11 well as short cycle breaths and prolonged cycle breaths (1, 8). Ineffective triggering was defined
12 as a simultaneous decrease in airway pressure and an increase in airflow without an assisted
13 breath (a wasted patient effort) (Figure 1). Double-triggered breaths occur when the ventilator
14 inspiratory time is shorter than the patient's inspiratory time in turn causing the incomplete
15 patient effort of the first cycle to trigger a second ventilator breath (Figure 2). Short cycle
16 breaths were defined as a cycle in which the inspiratory time is less than half the mean set
17 inspiratory time (Figure 3). Prolonged cycle breaths describe breaths in which the inspiratory
18 time is more than double the mean set inspiratory time (Figure 4). A unique asynchronous
19 breath type was identified during the study and labeled as ventilator breath stacking during
20 synchronized intermittent mandatory ventilation (SIMV). This ventilator breath stacking was
21 defined as breaths in which a mandatory breath was delivered during the inspiratory phase of a
22 spontaneous breath (with or without pressure support) (Figure 5).

1 Individual patient pressure, flow, and volume recorded waveforms were reviewed
2 simultaneously by two investigators blinded to demographic and outcome data. A third
3 investigator was utilized in circumstances of non-agreement with agreement between all
4 evaluators necessary before classifications as an asynchronous event occurred. The total breaths,
5 including non-asynchronous and asynchronous breaths (ineffective triggered, double-triggered,
6 short cycle, prolonged cycle and ventilator breath stacking) were calculated for each patient. The
7 asynchrony index (AI) was calculated by dividing the number of asynchronous breaths by the
8 number of asynchronous and non-asynchronous breaths (Asynchrony Index = Number of
9 Asynchronous Events /Total Number of Breaths.) Patients with an $AI \geq 10\%$ were defined as
10 asynchronous (1,8).

11 **Patient Characteristics**

12 Patient demographics including age, gender, mechanism of traumatic injury, Injury
13 Severity Score (ISS) and individual Chest Abbreviated Injury Scores (chest AIS), smoking
14 history and history of chronic obstructive pulmonary disease (COPD) were recorded (9).
15 Ventilator settings and arterial blood gas values prior to the initiation of the recording period
16 were noted. The assessment of sedation level and the presence of delirium was performed by
17 bedside ICU nurses versed in the utilization of the Richmond Agitation and Sedation Score
18 (RASS) and the Confusion Assessment Method in the ICU (CAM-ICU) in adherence to a
19 previously published ICU sedation protocol (10, 11, 12). The protocolized scoring of pain by the
20 visual analog score, sedation by the RASS and delirium by the CAM-ICU occurs on an every 8-
21 hour basis or when a change in behavior necessitates an intervention. Predetermined bolus
22 dosing of sedative and/or analgesic medications occurs first in response to objective score
23 changes with an increase in hourly rates of these same mediations if bolus therapy fails. The

1 treatment of delirium with haloperidol occurs only via bolus therapy though the frequency of
2 therapy may increase if subjective increases in delirium are present. Propofol and fentanyl are
3 utilized as the sedative and analgesic medications of choice within the first 72 hours after
4 intubation. Both RASS and CAM-ICU scores were recorded immediately prior to waveform
5 recording. The amount of fentanyl (mcg/kg), propofol (mg/kg) and haloperidol (mg) for the
6 previous 24 hours and 1 hour prior to recording were documented.

7 **Data Analysis**

8 The primary outcome was determined to be the number of ventilator days utilized
9 between patients with AI \geq 10% vs. those with AI $<$ 10%. Secondary outcomes of ICU length of
10 stay (LOS), hospital LOS, proportion of patients discharged home, and mortality were analyzed
11 between the AI groups.

12 Continuous data (non-normally distributed) were summarized using medians
13 [interquartile ranges], while categorical data were summarized using frequencies and percents.
14 Continuous data were compared between groups using Wilcoxon rank-sum tests. Categorical
15 data were compared using exact χ^2 chi-square tests. All tests were two-sided and $p \leq 0.05$ were
16 considered statistically significance. Analysis was carried out using SAS 9.2 (SAS Institute,
17 Inc., Carey, NC).

18

19 **RESULTS**

20 A total of 70 trauma patients were screened with 35 being ineligible (20 without
21 spontaneous respirations, 12 in which consent could not be obtained and 3 refused participation)

1 and 35 enrolled in the study (Figure 6). Patients were studied for 30 minutes each yielding a
2 total of 15,445 breath waveforms for analysis. None of the patients enrolled experienced
3 hemodynamic instability or required inotropic and/or vasopressor pharmacotherapy during
4 waveform collection. Twenty-six patients (74.3%) had an AI < 10%, and 9 (25.7%) had an AI ≥
5 10%. Demographics of the two cohorts are presented in Table 1. No statistical difference was
6 seen in median age, ISS, and chest AIS. There was no difference in the proportion of male
7 patients, the rate of penetrating injuries, history of smoking or COPD between the groups.
8 Arterial blood gas analyses between the cohorts lacked a difference in regards to pH, PaO₂/FIO₂
9 ratios, PaCO₂, and bicarbonate levels at the commencement of waveform analysis (Table 1).

10 When ventilator characteristics were analyzed, patients with an AI ≥ 10% had a
11 statistically greater proportion of synchronized intermittent mandatory ventilation (volume
12 control) with pressure support ventilation (SIMV + PSV) use (100% vs. 38.5%, $p = 0.0015$) with
13 a resultant higher median set rate of ventilator breaths per minute (12 [8, 14] vs. 0 [0, 6], $p <$
14 0.0001) and lower median rate of spontaneous breaths per minute (4 [3, 8] vs. 12 [9, 14], $p =$
15 0.02 ; Table 2). SIMV + PSV with a set respiratory rate of ≥ 10 breaths per minute was
16 significantly more prevalent in the AI ≥ 10% cohort (85.7% vs. 14.3%, $p = 0.02$).

17 The sedation levels, delirium rates, analgesic, sedative and haloperidol use were
18 evaluated between the cohorts. The proportion of patients with a RASS between +1 to -1 and
19 rate of positive CAM-ICU evaluations at the time of waveform collection was similar between
20 those with < 10% AI and ≥ 10% AI (Table 2). No difference was found between the groups for
21 the use of propofol (mg/kg), fentanyl (mcg/kg), and haloperidol (mg) in the previous hour and 24

1 hours before waveform recordings were performed. No haloperidol was provided to any patient
2 up to one hour immediately prior to recording.

3 Breath characteristics were analyzed for both AI cohorts (Table 3). The $\geq 10\%$ AI cohort
4 had 10,896 total breaths of which 1,184 were asynchronous while the $< 10\%$ AI group had a total
5 of 4,549 breaths, 235 were asynchronous. A statistically different pattern of asynchronous types
6 were seen between the groups ($p < 0.01$) with the $\geq 10\%$ AI cohort having more prolonged cycle
7 breaths (54.7% vs. 11.5%) and less ventilator breath stacking (26.2% vs. 54.9%) compared to
8 those without group asynchrony. Since ventilator breath stacking was not among previously
9 described asynchrony breath types, secondary statistical analysis was performed excluding these
10 breath types. With this breath exclusion, the proportion of asynchrony within the cohort did not
11 significantly differ.

12 Patients with an AI $\geq 10\%$ failed to demonstrate a significant outcome difference (Table
13 4). Ventilator days, ICU LOS, hospital LOS, the proportion of patients discharged home and
14 mortality were similar between the cohorts. The inclusion of long cycle and short cycle breaths
15 as an asynchronous subtype may have altered outcome interpretation therefore secondary
16 analyses were performed removing these episodes. Without the inclusion of these breaths, 29
17 patients were found to have $< 10\%$ AI and 6 patients had $\geq 10\%$ AI. Demographic and arterial
18 blood gas data were not statistically different between the AI groups. Ventilator settings and
19 proportion of those with SIMV with a ventilator rate ≥ 10 was not different between AI types
20 though the patients' respiratory rate became non-significant. The amount of fentanyl and
21 propofol, RASS and CAM-ICU scores were not statistically different as well. The proportion of
22 asynchronous breath types were statistically different ($p < 0.001$) between those with $< 10\%$ and
23 $\geq 10\%$ AI. Ventilator breath stacking was the majority of the asynchronous breaths in the $< 10\%$

1 cohort (59.7%) while missed/ineffective trigger breaths made up the majority in those with \geq
2 10% AI (43.8%). There was no statistically significant outcome difference demonstrated in
3 regards to ventilator days, ICU LOS, hospital LOS, proportion discharged home and mortality
4 seen between AI groups when long cycle and short cycle asynchronies were removed.

5

6 **DISCUSSION**

7 In this study, we investigated the complex interaction of mechanical ventilation with a
8 traumatically injured population so to clarify the presence, etiology and outcomes from patient-
9 ventilator asynchrony. We demonstrated that approximately a quarter of traumatically injured
10 patients exhibit ventilator asynchrony ($AI \geq 10\%$) and that these patients have an associated
11 greater utilization of SIMV + PSV with a set rate ≥ 10 breaths per minute. Unique to this work
12 is the finding of ventilator breath stacking, a mandatory breath delivered during the inspiratory
13 phase of a spontaneous breath, as an additional asynchronous breath type. Nonetheless, the
14 presence of asynchrony was not associated with prolonged mechanical ventilation, ICU LOS,
15 hospital LOS, or mortality in this population. It is noteworthy that while ineffective triggers are
16 the most commonly identified asynchrony in medical patients, missed triggers were relatively
17 uncommon in our study. This is likely due to the presence of chronic lung disease and air-
18 trapping in prior studies which complicate triggering in the medical patient. In a number of our
19 patients, intrinsic positive end expiratory pressure (PEEPi) was present, but in the absence of
20 dynamic hyperinflation and respiratory muscle dysfunction, did not result in missed triggers.

21 Patient-ventilator asynchrony is a common problem encountered in critical care
22 platforms and is associated with poor outcomes. Chao et al. demonstrated in 174 chronically

1 ventilator dependent patients that those with ineffective triggered breaths required a longer
2 duration of mechanical ventilation (70 vs. 33 days) and lower rate of eventual liberation (16% vs.
3 57%) (6). Further work by Thille et al. reaffirmed an association of prolonged mechanical
4 ventilation with an AI \geq 10% (median duration 25 vs. 9 days) as well as describing a higher rate
5 of tracheostomy (33% vs. 4%) (1). However, Chao et al analysis was performed in older
6 critically ill medical populations (69-75 years old) with 45% having the diagnosis of COPD
7 while the work of Thille et al was performed in ICU patients with a quarter having COPD.
8 Ineffective triggering/missed triggering breaths are the most common (85-88%) asynchronous
9 breaths recorded in previous medical studies (1,8). These breaths are associated with PEEPi
10 resulting from large tidal volumes and the continuation of mechanical inspiration during patient
11 defined expiration (3,4,13). Patients with COPD have a higher rate of ineffective triggering as
12 pressure support levels increase because higher patient generated pressures are required to
13 overcome the PEEPi associated with hyperinflation and shortened expiratory time
14 (3,6,7,14,15,16).

15 Little is known about patient-ventilator asynchrony in a traumatically injured patient
16 population. In comparison to a medically ill population, our study trauma population is younger
17 (median 47 years) with a resultant decreased rate of COPD (16.1%). Even with these inherent
18 differences, the proportion of asynchronous patients (25.7%) is comparable to previous studies
19 even in the presence of light sedation goals with delirium monitoring and treatment (1,8). Our
20 patient population exhibited unique proportions of asynchrony breath types. With a decreased
21 rate of COPD in this population, ineffective triggering occurred infrequently (10.1%) as a
22 proportion of asynchronous breaths while prolonged cycle breaths (54.7%) were the most
23 common asynchronous breath type in those with \geq 10% AI.

1 Upon further examination of the prolonged cycle breaths, we discovered that in the \geq
2 10% AI group, one patient had 75% of the total of this breath type. Additionally, this patient had
3 the highest set ventilator rate in SIMV than any other patient in the study which is an associated
4 risk factor for high asynchrony. In light of this finding, re-analysis of asynchronous breath type
5 proportions excluding this patient was performed. With this data excluded, the difference in
6 prolonged cycle breaths between the low ($< 10\%$ AI) and high asynchrony groups ($\geq 10\%$ AI)
7 was still statistically significant ($p < 0.01$) but only comprised 23.3% of the total asynchronous
8 breaths instead of 54.7% as shown in Table 3. Nonetheless, prolonged cycle breaths are still a
9 significant source of asynchronous breaths in the trauma patient population.

10 Unique to this study was the identification of what we termed ventilator breath stacking.
11 This occurs when a mandatory breath is delivered during the inspiratory phase of a spontaneous
12 breath. This phenomenon occurred during mandatory ventilator breaths in SIMV using the dual
13 control feature (Auto Flow) on the Drager XL ventilator. When this feature is activated, the
14 software measures dynamic respiratory system compliance on a breath-to-breath basis and
15 calculates the pressure required to reach the target tidal volume on the next mandatory breath.
16 Functionally, this mode is pressure limited but uses a feedback loop to target an inspired tidal
17 volume. The breath type is pressure controlled with decelerating flow. By definition, the
18 ventilator algorithm synchronizes the SIMV mandatory breaths with the patient's spontaneous
19 breaths to eliminate stacking a mandatory breath on top of a spontaneous one, resulting in a
20 delivered volume much larger than set. We observed that in the presence of Auto Flow,
21 synchronization of the mandatory breath and patient effort was not always obtained. Even
22 though the mandatory breath is delivered on top of the spontaneous breath, since the breaths are
23 pressure limited, over-inflation might be avoided. Though the delivered tidal volume is not

1 affected, this phenomenon increases inspiratory time which is often longer than neural
2 inspiratory time of spontaneous breaths and may result in asynchrony.

3 Our study has several limitations. Though the use of ventilator waveforms have been
4 previously validated, this study did not utilize esophageal pressure monitoring for the
5 determination of triggering events. As such, an inherent lack of sensitivity for detection may
6 exist. Even with a predetermined window (first 48 hours after intubation) for waveform
7 recording performed over 30 minutes, periods of asynchrony could be missed thus
8 underestimating the prevalence of this phenomenon in our population. Furthermore, recordings
9 were occurred during daytime hours and thus might have omitted asynchronous periods that
10 occurred during the night when interruptions in sleep by normal ICU care may account for
11 periods of agitation and/or delirium. Though not statistically significant, a trend towards RASS
12 scores outside of -1 to +1 was associated with those with a higher asynchrony index. The
13 smaller sample size of this study may have contributed to false conclusions in regards to sedation
14 differences between the groups. We used the Drager XL ventilator exclusively for this study due
15 to its compatibility with the recording software. We assume that other ICU ventilators would
16 produce similar patient-ventilator interactions using the same modes. Finally, this study was
17 conducted in a single institution with 35 trauma patients using various modes of mechanical
18 ventilation. Our study may not have the power to find an outcome difference between the two
19 groups, therefore a larger cohort of trauma patients with uniformity in regards to mechanical
20 ventilation mode over multiple institutions may support alternative findings. Future work may
21 focus on a larger population of trauma patients compared both internally and externally to both
22 surgical and non-surgical cohorts.

23

1 **CONCLUSIONS**

2 In this current work, patient-ventilator asynchrony is common in mechanically ventilated
3 traumatically injured patients. However, associations between groups with high asynchrony and
4 negative outcomes including ventilator days, ICU and hospital LOS, and mortality could not be
5 made. The most common ventilator mode associated with high asynchrony in our patient
6 population was SIMV with set respiratory rate ≥ 10 bpm. While ineffective efforts have
7 represented as much as 85% of asynchronous breaths in medical patients, this was not observed
8 in this cohort of surgical patients. Careful matching of the appropriate ventilator mode(s) and
9 settings to the patient's ventilatory needs may help minimize asynchrony and improve patient
10 comfort.

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20 **FIGURE LEGENDS**

1 **Figure 1.** Missed/ineffective trigger characterized by a decrease in airway pressure and an
2 increase in airflow without an assisted breath. Ventilator settings were PSV Δ 10 cm H₂O over 5
3 cm H₂O PEEP.

4
5 **Figure 2.** Double triggering is the result of the ventilator inspiratory time being shorter than the
6 patient's inspiratory time causing the incomplete patient effort of the first cycle to trigger a
7 second ventilator breath. Ventilator settings were SIMV + PSV Δ 10 cm H₂O over 5 cm H₂O
8 PEEP, with a set respiratory rate of 12 breaths/min. Mandatory breaths are adaptive pressure
9 breaths (autoflow).

10
11 **Figure 3.** Short cycle breaths have an inspiratory time that is less than half the mean inspiratory
12 time. Ventilator settings were PSV Δ 10 cm H₂O over 5 cm H₂O PEEP.

13
14 **Figure 4.** Prolonged cycle breaths occur when the ventilator's inspiratory time is more than
15 double the patient's inspiratory time. Ventilator settings were PSV Δ 10 cm H₂O over 5 cm H₂O
16 PEEP.

17
18 **Figure 5.** Ventilator breath stacking occurs when a mandatory breath was delivered during the
19 inspiratory phase of a spontaneous breath. Ventilator settings were SIMV + PSV (with
20 autoflow) Δ 10 cm H₂O over 5 cm H₂O PEEP with a set respiratory rate of 12 breaths/min.
21 Mandatory breaths are adaptive pressure breaths (autoflow).

22 **Figure 6.** Flow diagram of the enrollment process.

	< 10% AI (n = 26)	≥ 10% AI (n = 9)	p-Value
Demographics			
Age	45 (33, 59)	48 (40, 58)	0.79
Male	80.8%	66.7%	0.39
Penetrating Injury	30.8%	22.2%	0.62
ISS	21 (17, 29)	22 (21, 25)	0.82
Chest AIS	2.5 (0, 8)	2 (0, 3)	0.63
History of Smoking	43.5%	16.7%	0.23
History of COPD	16.7%	14.3%	0.88
Arterial Blood Gas			
pH	7.38 (7.35, 7.42)	7.41 (7.32, 7.44)	0.97
PaO ₂ /FiO ₂ Ratio	283 (254, 350)	395 (200, 468)	0.57
PaCO ₂	43 (38, 47)	38 (35, 43)	0.14
Bicarbonate	25 (22, 27)	23 (23, 26)	0.34

Table 1. Demographics and Arterial Blood Gas on Evaluation

All values medians (interquartile ranges) unless otherwise specified, $p < 0.05$ considered significant AI, asynchrony index; ISS, Injury Severity Scale; AIS, Abbreviated Injury Scale; COPD, Chronic Obstructive Pulmonary Disease

	< 10% AI (n = 26)	≥ 10%AI (n=9)	p-Value
Ventilator			
SIMV + PSV	38.5%	100%	0.0015
Tidal Volume (mL)	600 (550, 600)	550 (550, 600)	0.63
Inspiratory Time (sec)	1.15 (0.9, 1.25)	1.2 (0.9, 1.2)	0.86
PIP (cm H ₂ O)	22 (16.5, 25)	19 (17, 26)	0.94
Pressure Support (cm H ₂ O)	10 (10, 10)	10 (10, 10)	0.48
PEEP (cm H ₂ O)	5 (5, 8)	5 (5, 5)	0.46
Ventilator Rate (BPM)	0 (0, 6)	12 (8, 14)	< 0.0001
Patient Rate (BPM)	12 (9, 14)	4 (3, 8)	0.007
SIMV with Vent Rate ≥ 10 (BPM)	14.3%	85.7%	0.02
Sedation/Delirium			
Fentanyl previous hour (mcg/kg)	1.01 (0.45, 1.35)	1.00 (0, 1.25)	0.51
Fentanyl previous 24 hours (mcg/kg)	13.8 (6.8, 29.7)	16.0 (2.9, 21.2)	0.61
Propofol previous hour (mg/kg)	1.35 (0, 2.4)	0.30 (0, 1.5)	0.29
Propofol previous 24 hours (mg/kg)	20.5 (5.2, 41.6)	10.6 (9.5, 20.9)	0.35
RASS of +1 to -1	61.5%	33.3%	0.31
CAM-ICU Positive	33.3%	40.0%	0.78

Table 2. Ventilator Settings, Sedation and Rate of Delirium on Evaluation

All values medians (inter-quartile ranges) unless otherwise specified, $p < 0.05$ considered significant
 AI, asynchrony index; SIMV, synchronized intermittent mandatory ventilation; PSV, pressure support ventilation;
 PIP, peak inspiratory pressure
 BPM, breaths per minute; RASS, Richmond Agitation Sedation Scale; CAM-ICU, Confusion Assessment Method
 for the ICU

	< 10% AI n = 26	≥ 10% AI (n = 9)	p-Value
Total Number of Breaths	4549	10,896	
Total Number of Asynchronous Breaths	235	1,184	
Asynchronous Breath Type			< 0.01
Missed/Ineffective Trigger	32 (13.6%)	120 (10.1%)	
Double Trigger	41 (17.4%)	107 (9.0%)	
Short Cycle	6 (2.6%)	0 (0.0%)	
Long Cycle	27 (11.5%)	647 (54.7%)	
Ventilator Breath Stacking	129 (54.9%)	310 (26.2%)	

Table 3. Proportion of asynchronous breath types by AI cohorts

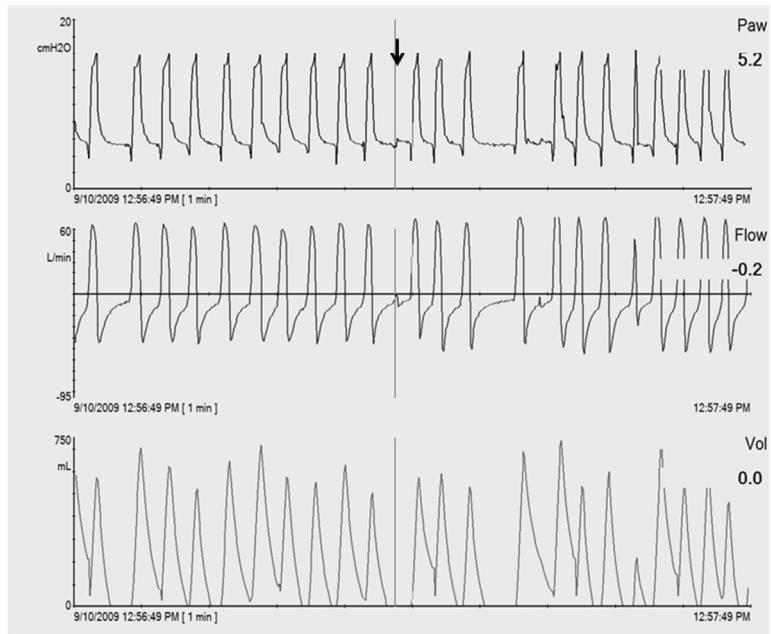
$p < 0.05$ considered significant

AI = asynchrony index

Outcomes			
Ventilator Days	7 (3, 14)	9 (4, 22)	0.42
ICU LOS (days)	11 (6, 18)	13 (8, 26)	0.28
Hospital LOS (days)	17 (11, 24)	22 (10, 27)	0.46
Discharged Home	38.5%	11.1%	0.13
Mortality	3.9%	11.1%	0.42

Table 4. Outcomes

All values medians (inter-quartile ranges) unless otherwise specified, $p < 0.05$ considered significant
AI, asynchrony index; LOS, length of stay



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Figure 2

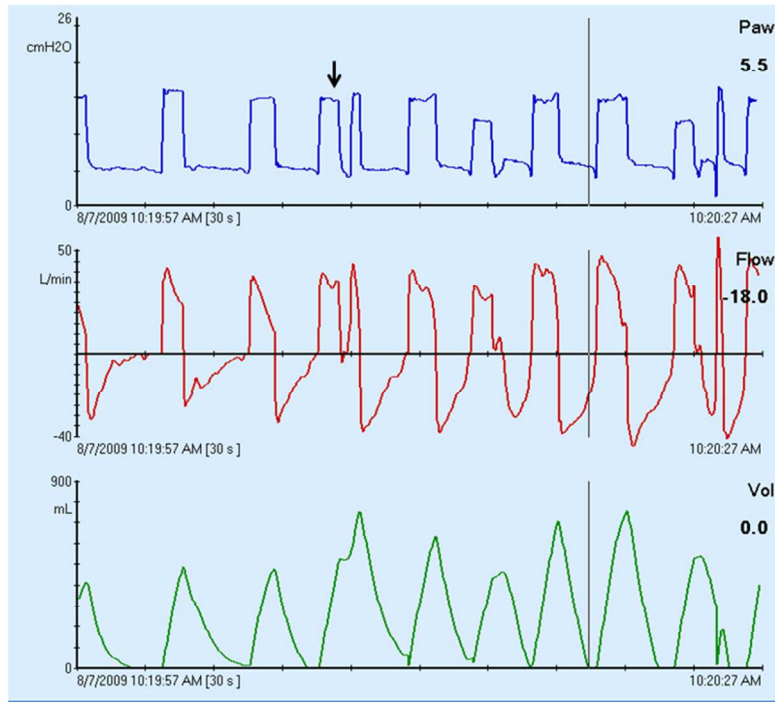
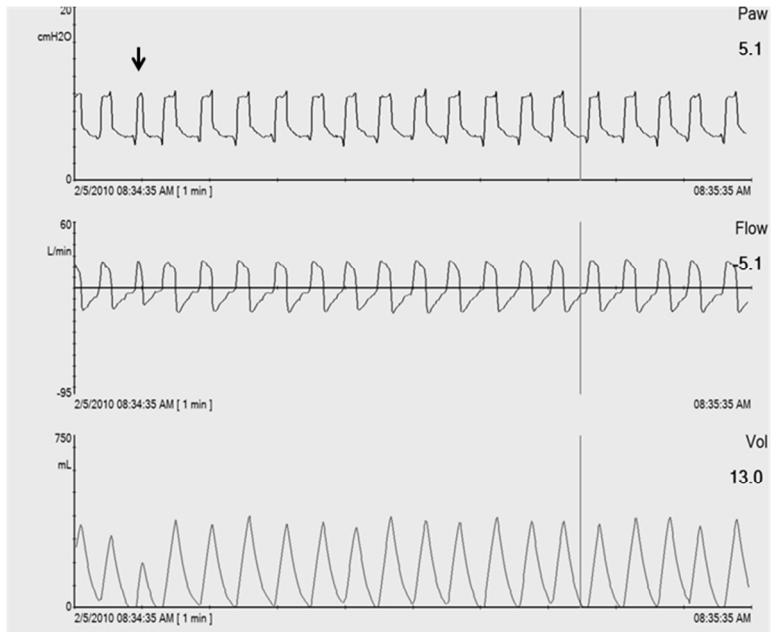
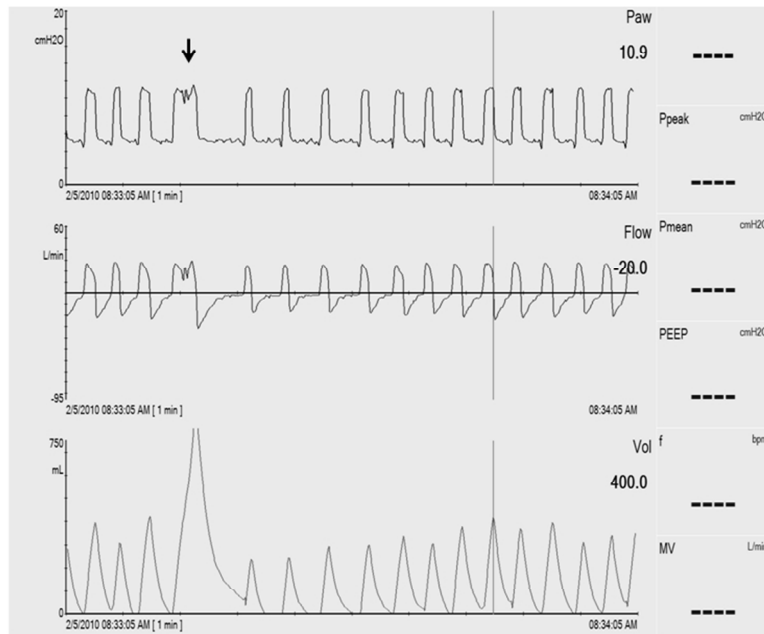


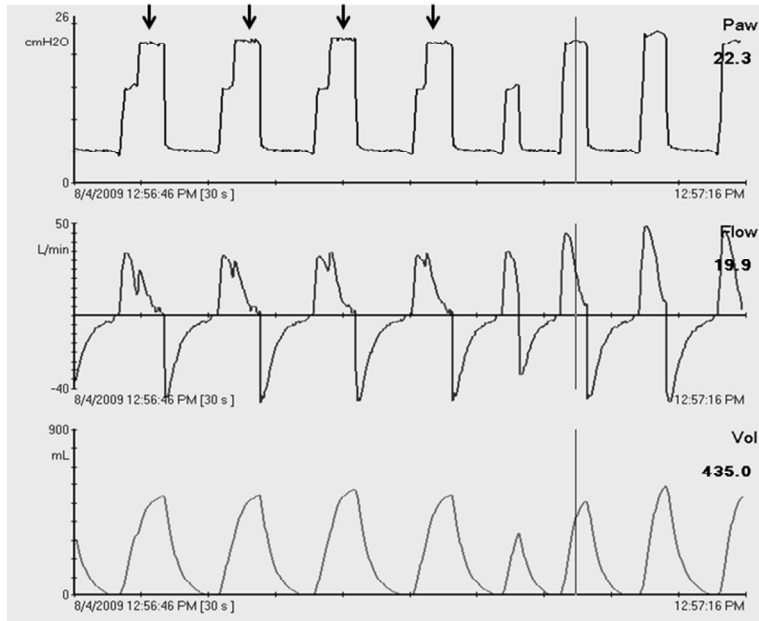
Figure 2
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