

Supplemental Table 1: Emory Protocol Compliance Score

NED Compliance (mcg/kg/min) <sup>a</sup>		
	NED $\leq$ 0.3 at time of initiation of Ang II	1
	NED 0.31 to 0.4 at time of initiation of Ang II	2
	NED 0.41 to 0.5 at time of initiation of Ang II	3
	NED $>$ 0.5 at time of initiation of Ang II	4
	NED $<$ 0.1 at time of initiation of Ang II	5
Vasopressin Compliance (U/min)		
	Vaso $\geq$ 0.06 at time of initiation of Ang II	1
	Vaso not started or $<$ 0.06 at time of initiation of Ang II	2
MB Compliance <sup>b</sup>		
	MB used after Vaso and before Ang II, or contraindicated	1
	MB not used after Vaso and before Ang II, and no contraindications	2
High Output Compliance (CI or ScvO <sub>2</sub> /CVP criteria) <sup>c</sup>		
	High output documented before Ang II started?	1
	High output documented after Ang II started?	2
	High output not documented?	3
	Low output documented?	4
Cumulative Compliance Score		
	Compliant	6 or less
	Moderately Compliant	7-8
	Poorly Compliant	$>$ 8

The Emory Protocol cumulative compliance score is calculated by adding the scores for each category (NED compliance, vasopressin compliance, MB compliance and high output compliance). Scores may range from 4 to 13. A cumulative compliance score of 6 or less is considered compliant, 7-8 as moderately compliant, and greater than 8 as non-compliant.

<sup>a</sup>Compliance is graded as best score between start time and order time on Ang II.

<sup>b</sup>Contraindications to MB use include antecedent use of selective serotonin release uptake inhibitors, serotonin-norepinephrine release uptake inhibitors, or monoamine oxidase inhibitors, or documentation of G6PD deficiency. <sup>c</sup>High output is determined based on either a CI $>$ 2.3 or ScvO<sub>2</sub> $\geq$ 70% + CVP $>$  8 cm H<sub>2</sub>O. Ang II, angiotensin II; CI, cardiac index; CVP, central venous pressure; MB, methylene blue; mcg, micrograms; ScvO<sub>2</sub>, mixed venous oxygen saturation; kg, kilograms; min, minute; NED, norepinephrine equivalence dose; U, units; Vaso, vasopressin.

Supplemental Table 2: Regression Model A of Unbalanced Variables for ICU Mortality

Variable	B	95% C.I.	<i>P</i>
Serum Lactate prior to Ang II	0.089	0.943 to 1.453	0.24
SOFA	0.180	0.987 to 1.127	0.07
Use of Mechanical Circulatory Support	-0.468	0.168 to 2.336	0.49

Cumulative Compliance Score	0.446	1.061 to 2.300	0.02
Cardiopulmonary Bypass Time	0.006	0.998 to 1.015	0.13
Chronic Lung Disease	0.977	0.709 to 9.949	0.15
Constant	-8.076		0.000

ICU, intensive care unit; Ang II, angiotensin II; SOFA, Sequential Organ Failure Assessment. Use of mechanical support and presence of chronic lung disease were considered categorical variables, with the other variables treated as continuous.

Supplemental Table 3: Regression Model B of Known Predictors for Mortality and Cumulative Compliance Score Against ICU Mortality

Variable	B	95% C.I.	<i>P</i>
Serum Lactate prior to Ang II	0.133	1.000 to 1.304	0.05
SOFA Score	0.150	0.974 to 1.386	0.10
Cumulative Compliance score	0.388	1.045 to 2.081	0.03
NE Dose at start of Ang II	-0.045	0.021 to 44.248	0.98
Constant	-5.715		0.001

ICU, intensive care unit; Ang II, angiotensin II; SOFA, Sequential Organ Failure Assessment; NE, norepinephrine equivalence. All variables were treated as continuous.

Supplemental Table 4: Regression Model C of Statistically significant Variables from Model A and B

Variable	B	95% C.I.	<i>P</i>
SOFA	0.182	0.988 to 1.455	0.07
Cumulative Compliance Score	0.443	1.065 to 2.278	0.02
Cardiopulmonary Bypass Time	0.007	0.999 to 1.015	0.08
Chronic Lung Disease	0.942	0.679 to 9.679	0.17
Serum Lactate prior to Ang II	0.091	0.944 to 1.269	0.23
Constant	-8.407		0.000

SOFA, Sequential Organ Failure Assessment. Both variables treated as continuous.

Supplemental Table 5: Regression of Individual Compliance Scores for ICU Mortality

Variable	B	95% C.I.	<i>P</i>
NE Dose Compliance Score	0.024	0.655 to 1.601	0.92
Vaso Dose Compliance Score	-1.735	0.024 to 1.300	0.09
Cardiac Output Compliance Score	0.713	1.205 to 3.454	0.008
MB Compliance Score	-0.282	0.252 to 2.255	0.61
Constant	-0.545		0.64

ICU, intensive care unit; NE, norepinephrine equivalence; Vaso, vasopressin. Vaso Dose Compliance Score was considered a categorical variable, with the other two variables treated as continuous.