

Angiotensin II v kardiochirurgii

Komu a kdy?

Hynek Říha

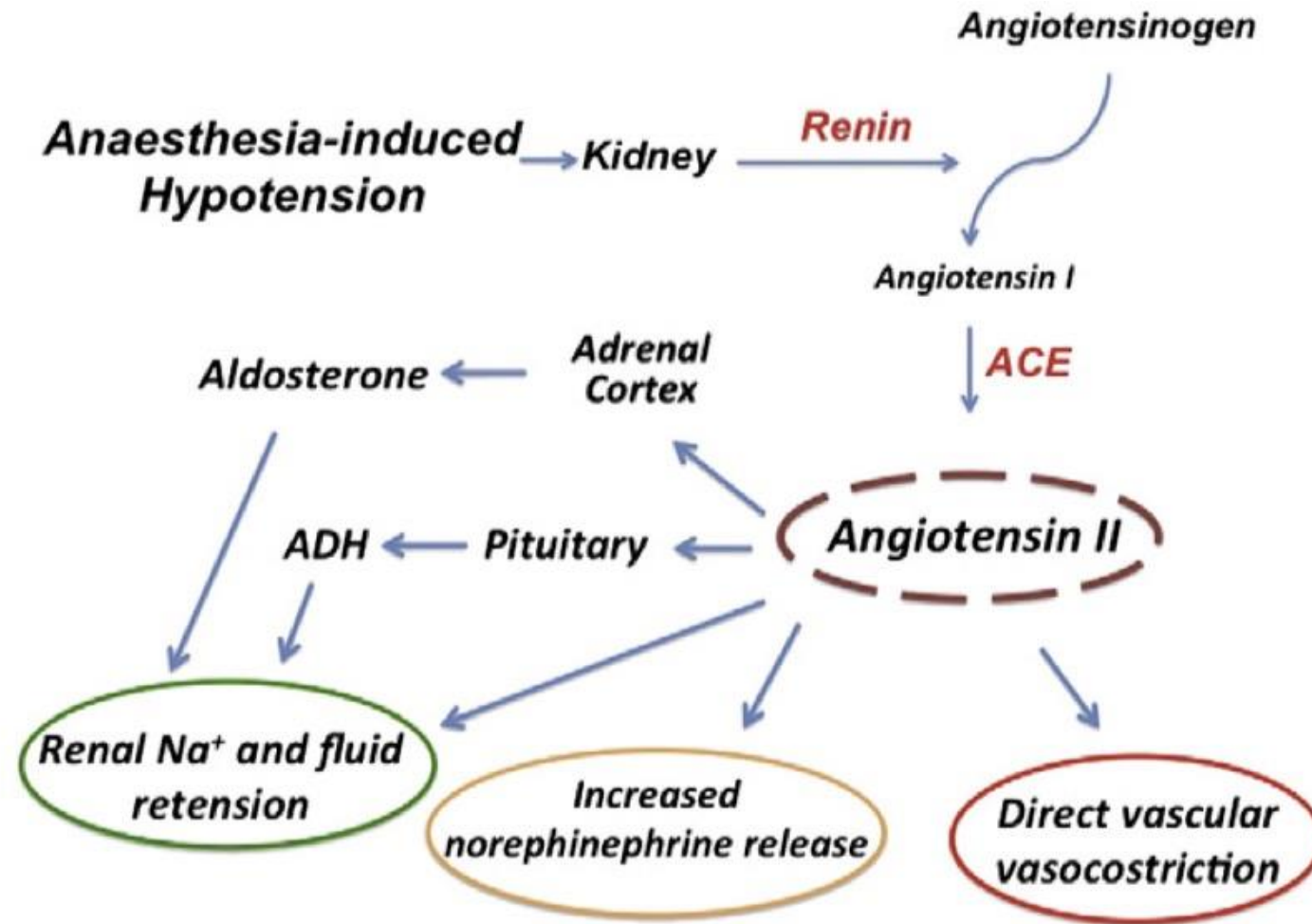
Klinika anesteziologie a resuscitace
Kardiocentrum IKEM, Praha



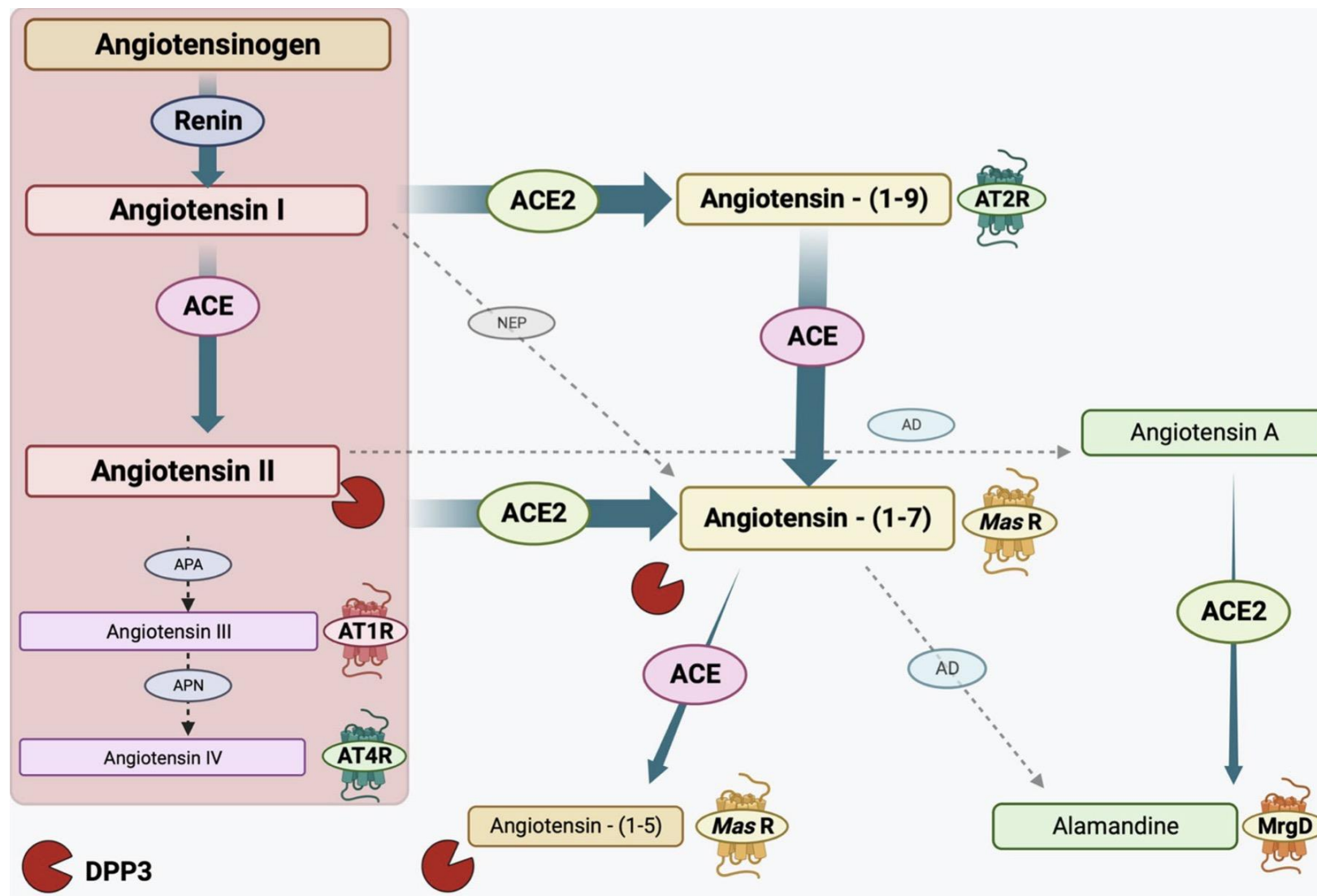
Deklarace konfliktu zájmů

- Medis
- ✓ 1x edukační přednáška 6/2025

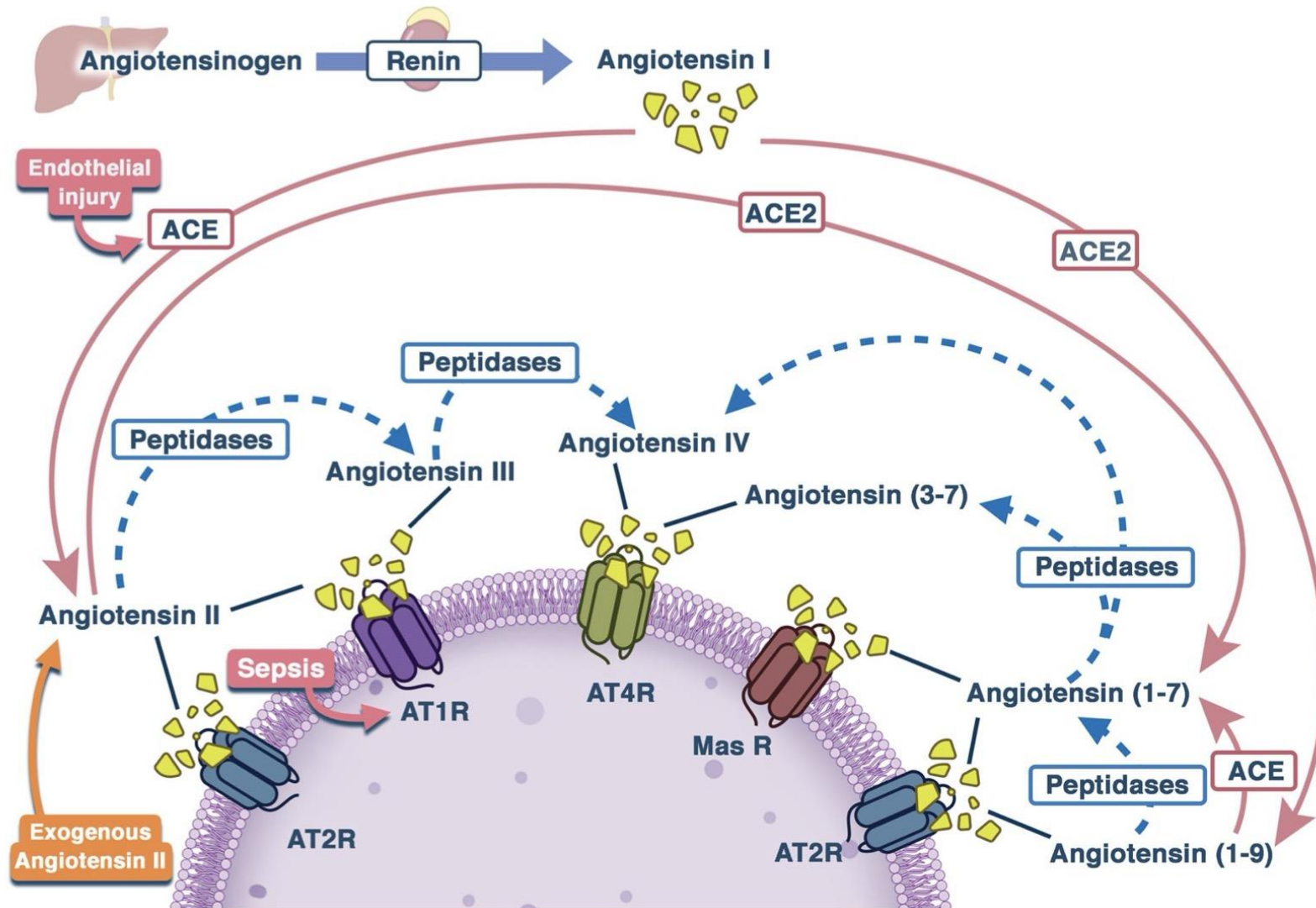
System renin – angiotenzin – aldosteron (RAAS)



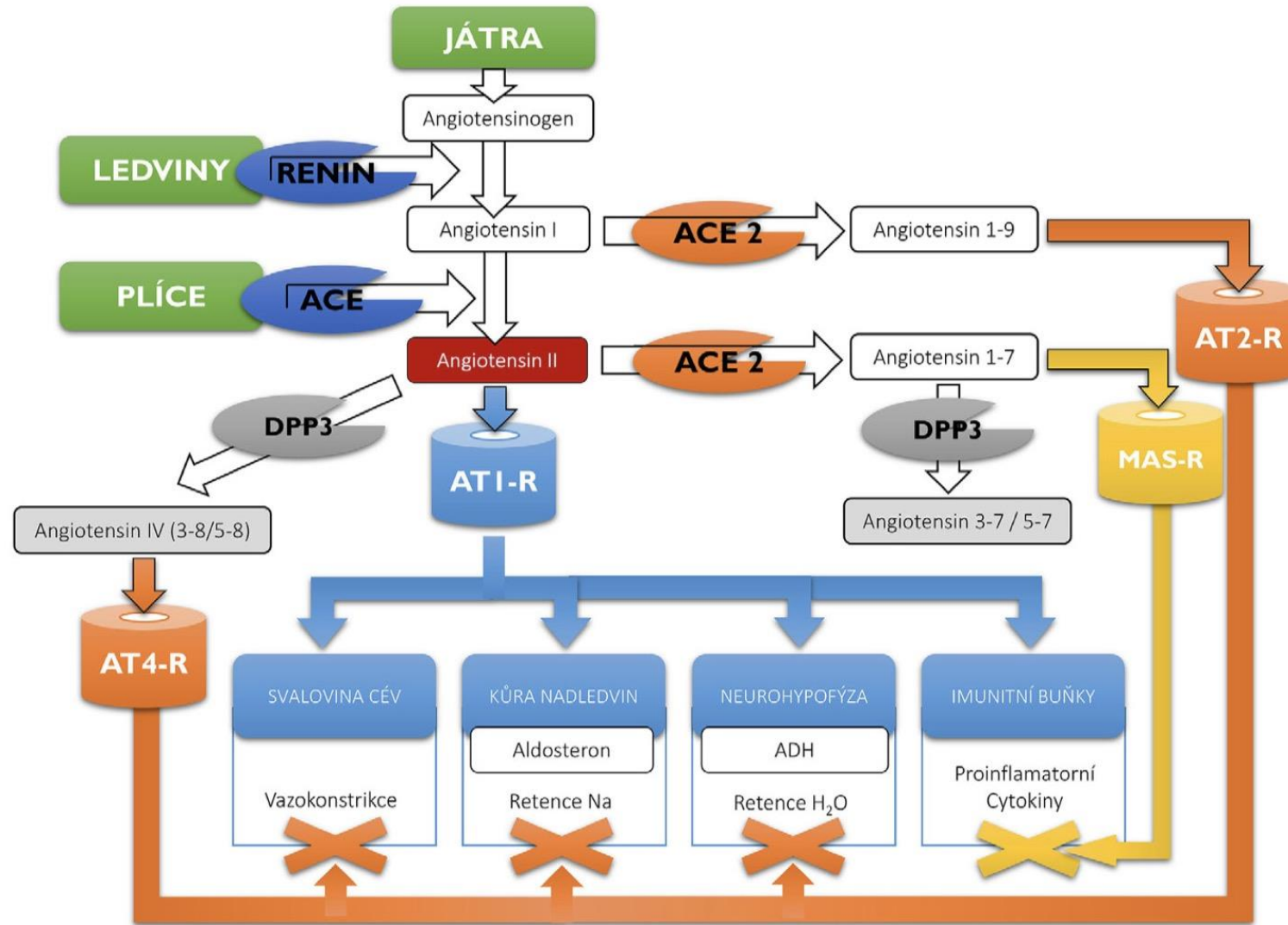
RAAS: klasická a alternativní cesta



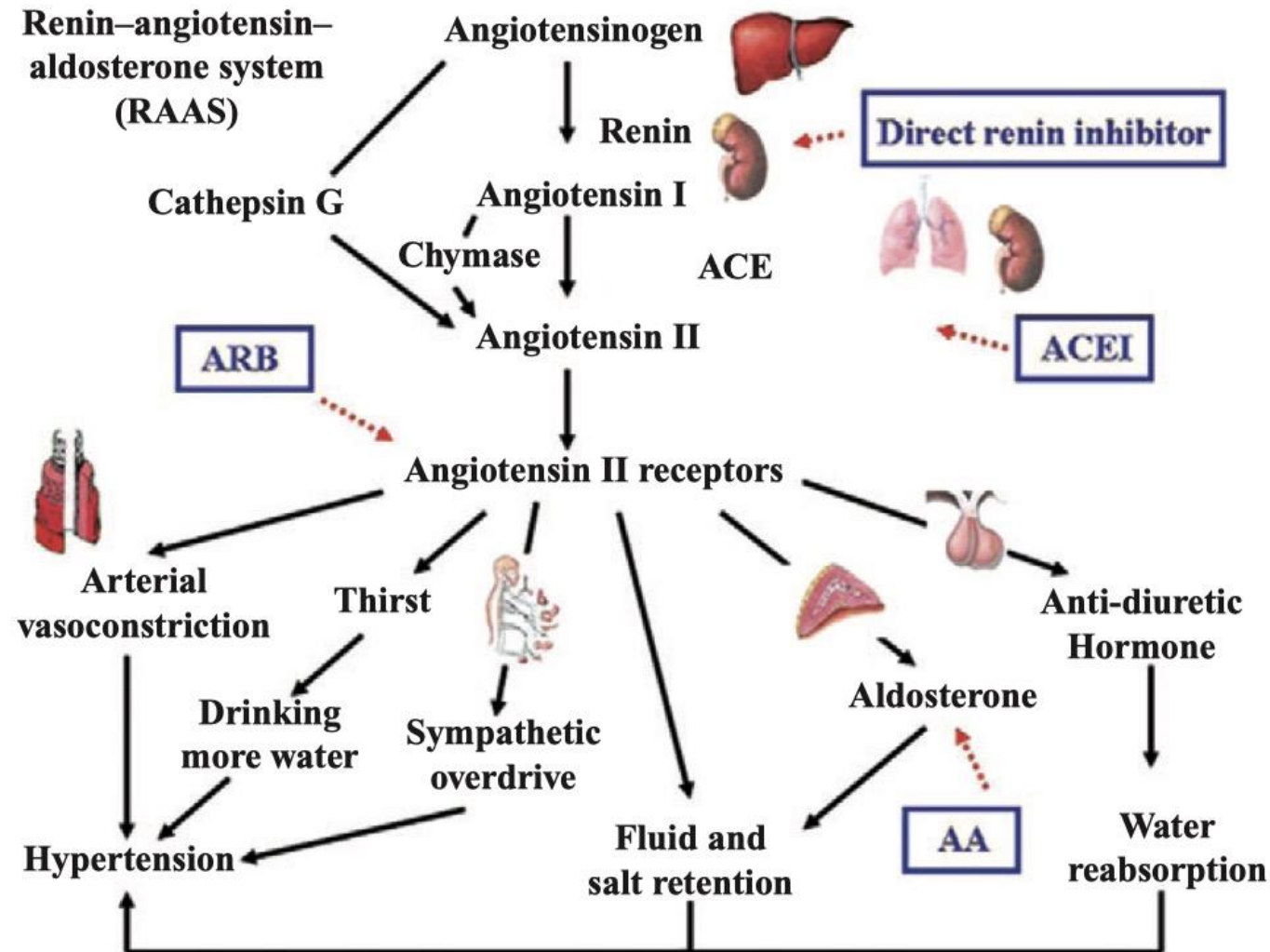
RAAS: komplexní systém



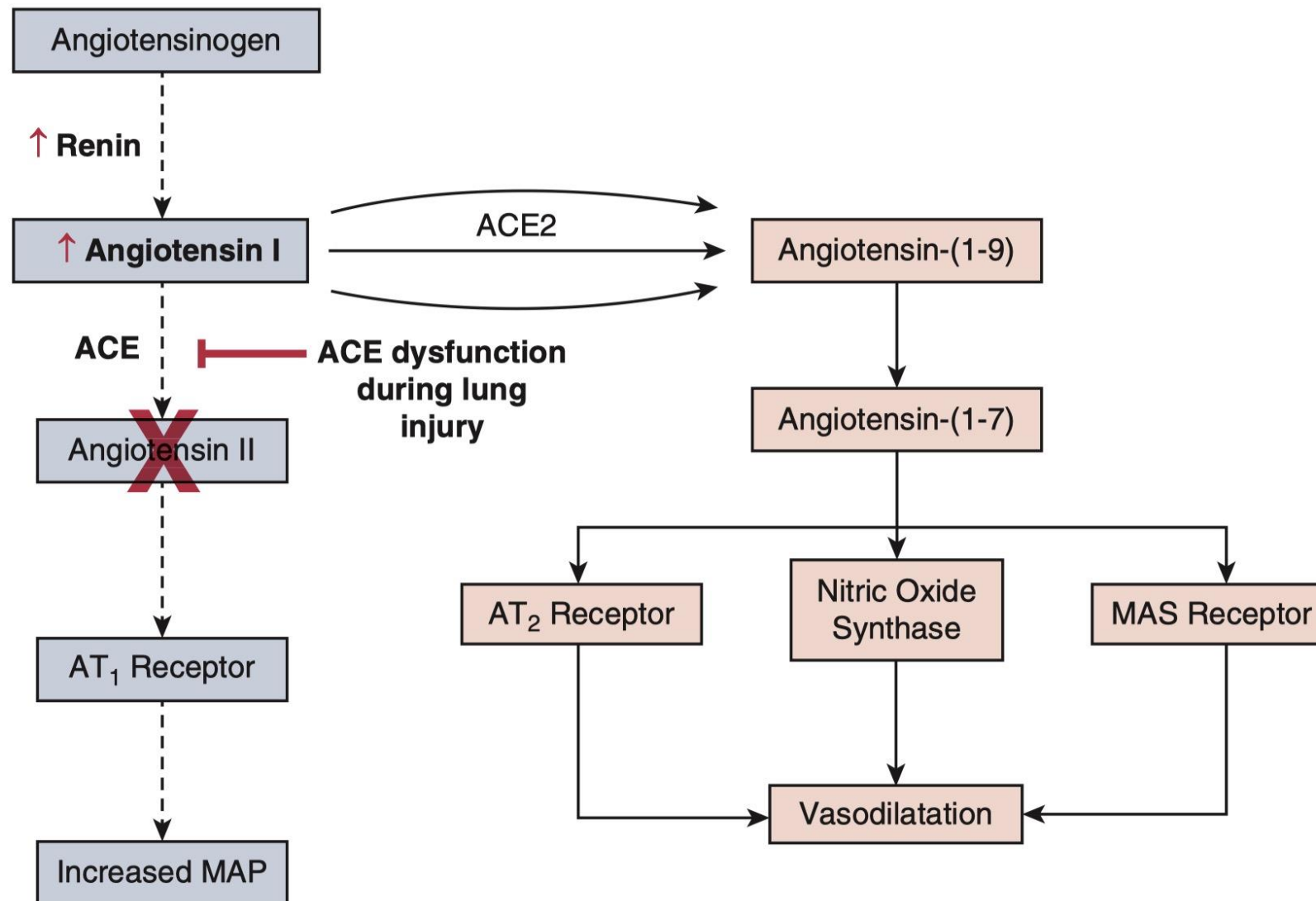
RAAS: komplexní systém



RAAS: ACE inhibitory a sartany



RAAS: KCH výkony (mimotělní oběh) a V-A ECMO



Vazoplegie / vazoplegický syndrom / vazoplegický šok

- Stav abnormálně nízké systémové cévní rezistence (SVR), který se projevuje jako výrazná hypotenze anebo nutnost podávat farmaka k zvrácení hypotenze (nekontrolovaná vazodilatace)
 - Srdeční výdej je normální anebo zvýšený
 - Odpověď cév na podávané vazopresory
- Hypoperfuze orgánových systémů
- Chybí jasná a přesná, klinicky založená, definice

Vazoplegie u kardiochirurgických pacientů

PATIENT-RELATED RISK FACTORS	Anemia, CKD, ESRD, HFrEF, recent MI (<5 days), hypothyroidism, older age, male sex, BMI>35 kg/m ² , BSA>1.9 m ² , high comorbidity/risk score (e.g. EUROSCORE, INTERMACS)
PREOPERATIVE RISK FACTORS	Anemia; diuretic, vasopressor, and/or heparin use
INTRAOPERATIVE RISK FACTORS	Anemia, transfusion of blood products, aortic cross clamp time, CPB time, total cardioplegia volume, nadir temperature



Definice vazoplegie u kardiochirurgických pacientů

Parameters and Thresholds Used in Published Definitions

Vasodilation Criterion	Hemodynamic Criterion	Vasoactive Drugs	Timeframe	Other Criteria
MAP $\leq$ 50-80 mmHg	CI $\geq$ 2.0-3.5 L $\cdot$ min ⁻¹ m ⁻²	Type: Norepinephrine Vasopressin Epinephrine Dobutamine Phenylephrine Methylene blue Epinephrine Vitamin B ₁₂ Metaraminol	Onset: <30 minutes to 4 days after CPB	HR > 80/min
SVR $\leq$ 500-900 dynes $\cdot$ seconds $\cdot$ cm ⁻⁵	CO >4-5 L $\cdot$ min ⁻¹	Dosage or NEE variations	Duration: single reading up to > 24 hours	Mixed/central venous oxygen saturation >60%-70%
SVR index $\leq$ 600-1800 dynes $\cdot$ seconds $\cdot$ cm ⁻⁵ m ⁻²	LVEF >50%-55%	Combination of one or more agents		Oxygen extraction <35%
SBP <70-90 mmHg	CVP <5 or >8 mmHg			Bicarbonate levels < 19-20 mEq/L
	PCWP <10 mmHg			

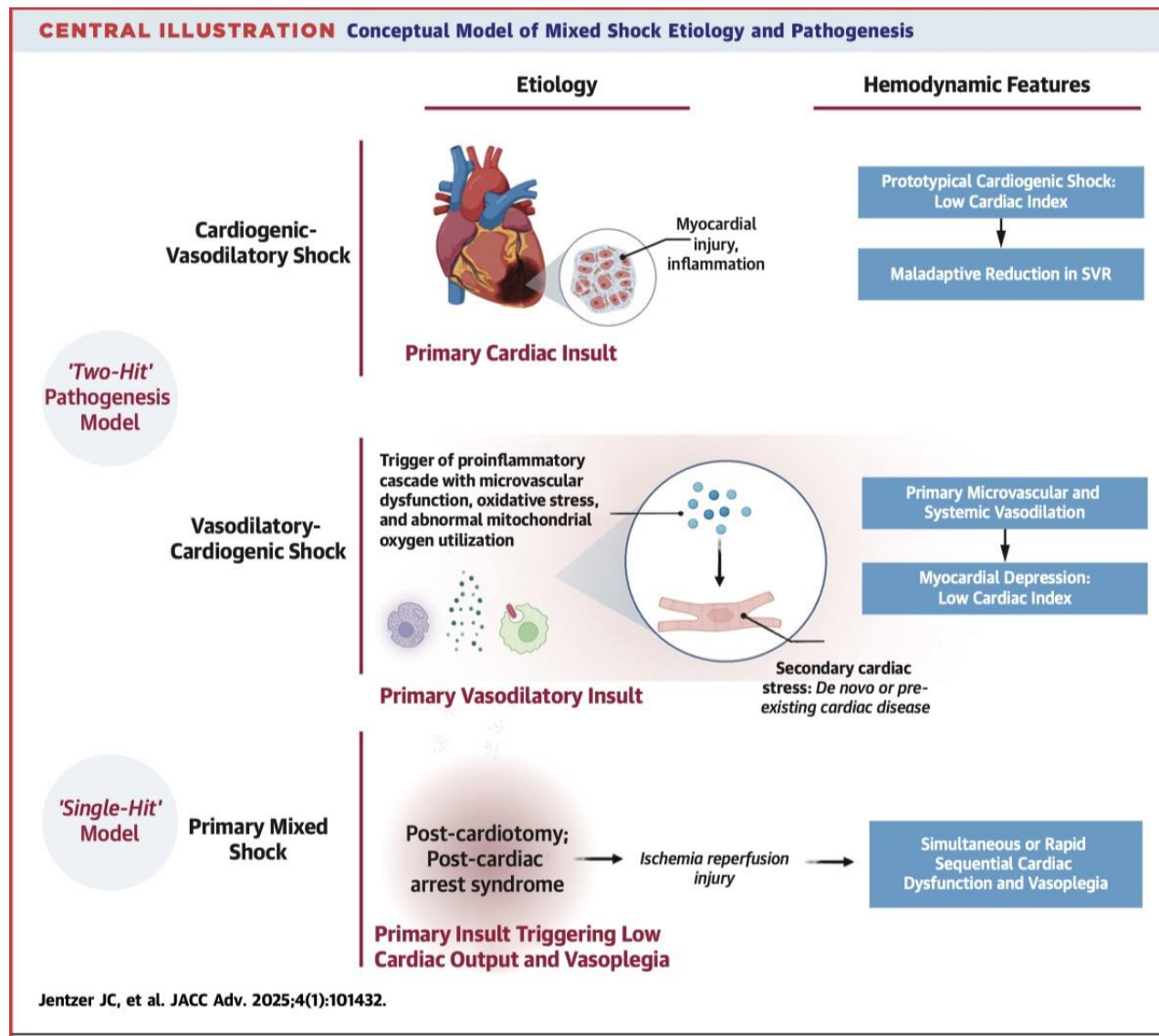
Papazisi O et al. JCVA 2025;39:1451

Vasoplegic Syndrome Consensus Definition Set by the Members of the Committee on Cardiovascular and Thoracic Anesthesia of the American Society of Anesthesiologists

Parameter	Suggestive of Vasoplegia
Mean arterial pressure	<65 mmHg
Cardiac index	$\geq$ 2.2 L/min/m ²
Systemic vascular resistance	<800 dynes $\cdot$ sec $\cdot$ cm ⁻⁵
Hemodynamic support	$\geq$ 0.2 μ g/kg/min norepinephrine equivalents
Rule out hypovolemia, intracardiac etiology, and adrenal insufficiency	

Polyak P et al. JCVA 2025;39:1815

Kombinovaný cirkulační šok



Fyziologie a patofyziologie systémové cévní rezistence

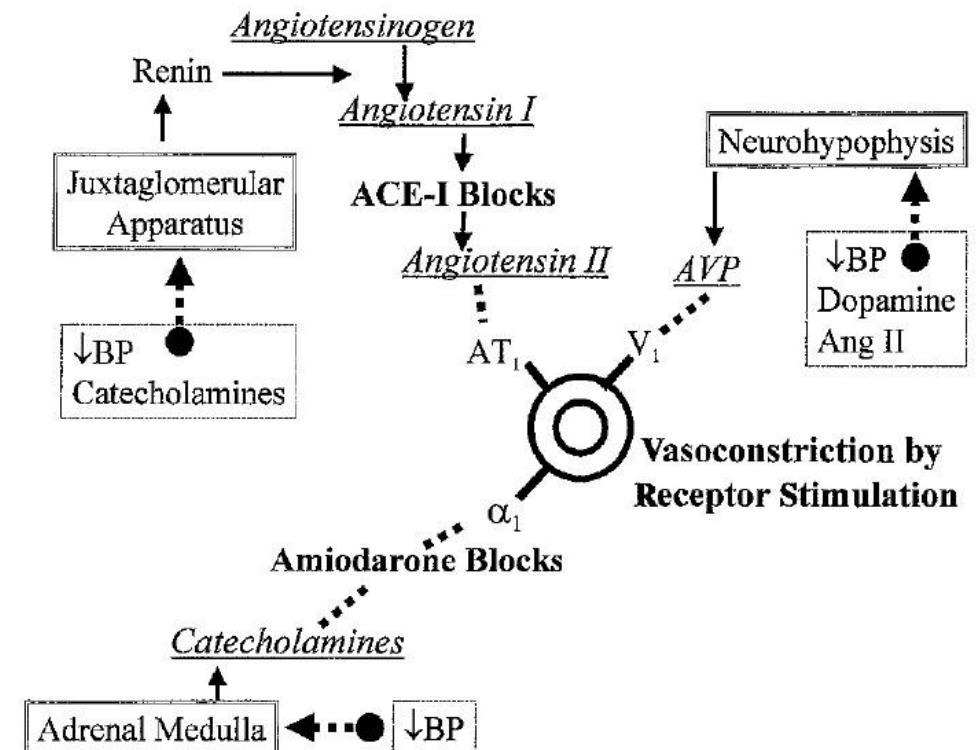
- Cévní tonus regulován intracelulární koncentrací Ca^{2+}

➤ Intrinsické regulátory

- ✓ NO, prostacyclin, endotelin, serotonin; acidóza, hypoxie ...

➤ Extrinsické regulátory

- ✓ Receptory VNS (sympatikus)
- ✓ Receptory pro vazopresin
- ✓ Receptory pro angiotenzin II



Farmakologické ovlivnění vazoplegie

Agent	MOA	Dose
Norepinephrine	Significant α_1 , α_2 agonism Moderate β_1 agonism	0.01–3 $\mu\text{g/kg/min}$
Epinephrine	Significant α_1 , α_2 agonism Significant β_1 agonism	0.01–1 $\mu\text{g/kg/min}$
Phenylephrine	Significant α_1 , α_2 agonism No effect on β_1	0.1–5 $\mu\text{g/kg/min}$
Dopamine	Dose dependent adrenergic agonism α_1 agonism as dose increases	1–20 $\mu\text{g/kg/min}$
Vasopressin	Repletion of vasopressin in ADH depleted state V1 agonism	0.01–0.1 U/min
Vitamin C	Cofactor for catecholamine synthesis	1.5 g every 6 h
Thiamine	Cofactor of lactate dehydrogenase (increase in lactate clearance)	100 mg every 6 h
Hydrocortisone	Aids in vitamin C metabolism Repletion of glucocorticoid and mineralocorticoid activity in cortisol depleted state Inhibition of pro-inflammatory cytokines	50 mg every 6 h
Methylene blue	Inhibition of guanylyl cyclase and inducible endothelial NO synthase	1–2 mg/kg
Hydroxocobalamin	Inhibition of NO directly and inducible endothelial NO synthase Inhibition of hydrogen sulfide	5 g
Angiotensin II	AT1 agonism Stimulation of aldosterone release Increase in ADH synthesis	10–40 ng/kg/min

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Angiotensin II for the Treatment of Vasodilatory Shock

Ashish Khanna, M.D., Shane W. English, M.D., Xueyuan S. Wang, M.D., Kealy Ham, M.D., James Tumlin, M.D., Harold Szerlip, M.D., Laurence W. Busse, M.D., Laith Altaweel, M.D., Timothy E. Albertson, M.D., M.P.H., Ph.D., Caleb Mackey, M.D., Michael T. McCurdy, M.D., David W. Boldt, M.D., Stefan Chock, M.D., Paul J. Young, M.B., Ch.B., Ph.D., Kenneth Krell, M.D., Richard G. Wunderink, M.D., Marlies Ostermann, M.D., Ph.D., Raghavan Murugan, M.D., Michelle N. Gong, M.D., Rakshit Panwar, M.D., Johanna Hästbacka, M.D., Ph.D., Raphael Favory, M.D., Ph.D., Balasubramanian Venkatesh, M.D., B. Taylor Thompson, M.D., Rinaldo Bellomo, M.D., Jeffrey Jensen, B.S., Stew Kroll, M.A., Lakhmir S. Chawla, M.D., George F. Tidmarsh, M.D., Ph.D., and Adam M. Deane, M.D., for the ATHOS-3 Investigators*

Studie ATHOS-3: uspořádání

- RCT: AT II vs. placebo (163 vs. 158 pacientů)
 - ✓ Vazodilatační šok s dávkou noradrenalinu $>0,2$ ug/kg/min
 - ✓ Sepse, pankreatitida, pooperační vazoplegie, multifaktoriální stav
- Primární výsledný ukazatel
 - ✓ MAP po 3 h od zahájení infuze
 - ✓ Vzestup MAP o ≥ 10 mmHg anebo MAP ≥ 75 mmHg
 - ✓ Bez zvýšení dávky ostatních vazopresorů
- Sekundární výsledné ukazatele
 - ✓ SOFA (kardiovaskulární, celkové)
- Další výsledné ukazatele (mortalita 7/28 dní)

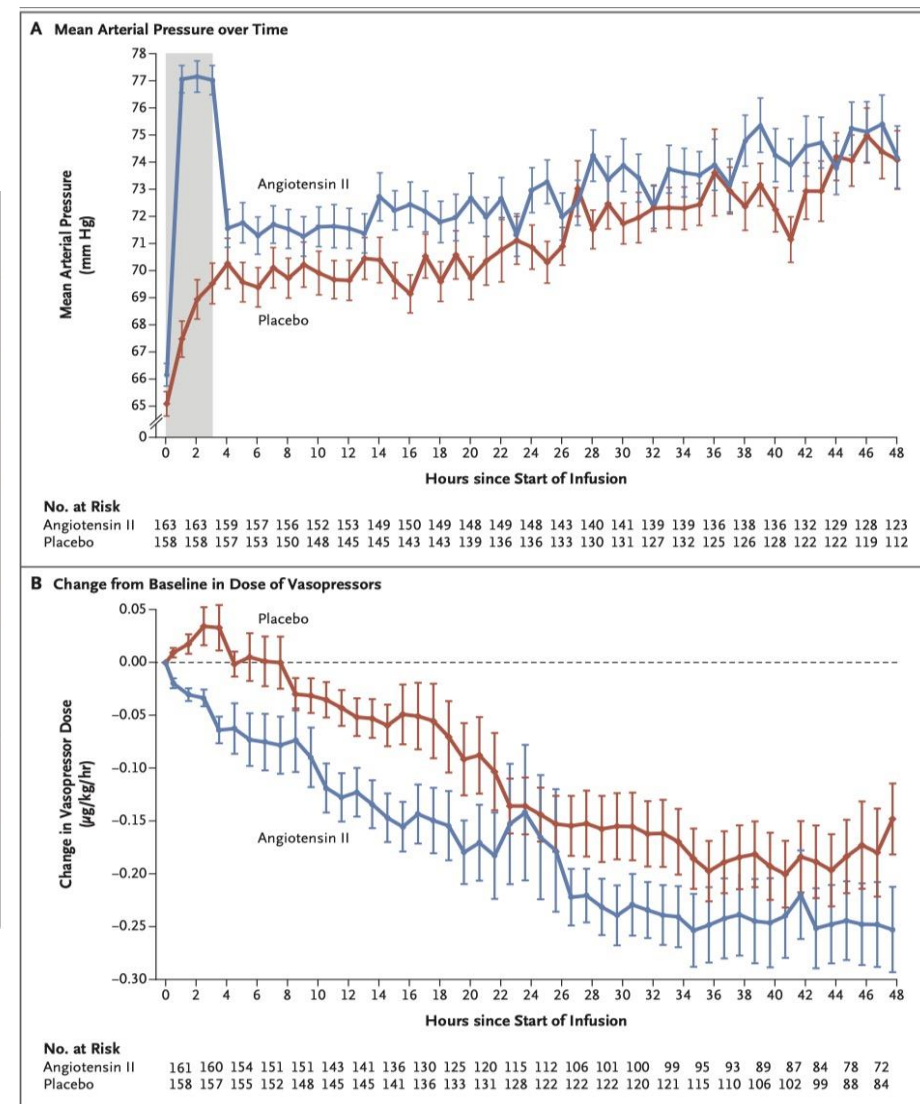
Studie ATHOS-3: výchozí charakteristika

Mean arterial pressure			
Median (IQR) — mm Hg	66.3 (63.7–69.0)	66.3 (63.0–68.3)	66.3 (63.7–68.7)
<65 mm Hg — no. (%)	52 (31.9)	50 (31.6)	102 (31.8)
APACHE II score‡			
Median (IQR)	27 (22–33)	29 (22–34)	28 (22–33)
Distribution — no. (%)			
≤30	105 (64.4)	93 (58.9)	198 (61.7)
31–40	50 (30.7)	54 (34.2)	104 (32.4)
≥41	8 (4.9)	11 (7.0)	19 (5.9)
Albumin			
Median (IQR) — g/dl	2.2 (1.8–2.7)	2.4 (1.9–2.8)	2.3 (1.8–2.7)
<2.5 g/dl — no./total no. (%)	103/154 (66.9)	89/156 (57.1)	192/310 (61.9)
ScvO ₂			
Median (IQR) — %	76.9 (73.0–82.8)	77.0 (72.5–82.0)	77.0 (72.9–82.2)
Data missing — no.	43	41	84
Central venous pressure			
Median (IQR) — mm Hg	13 (10–15)	12 (10–16)	12 (10–15)
Data missing — no.	37	35	72
Cardiac index			
Median (IQR) — liters/min/m ²	3.0 (2.6–3.8)	3.2 (2.7–3.9)	3.1 (2.6–3.8)
Data missing — no.	94	85	179
Median MELD score (IQR)§	21 (15–25)	23 (17–26)	22 (16–26)
Cause of shock — no. (%)			
Sepsis	127 (77.9)	132 (83.5)	259 (80.7)
Other, potentially sepsis	20 (12.3)	11 (7.0)	31 (9.7)
Pancreatitis	0	2 (1.3)	2 (0.6)
Postoperative vasoplegia	10 (6.1)	9 (5.7)	19 (5.9)
Multifactorial	6 (3.7)	4 (2.5)	10 (3.1)
Exposure to ACE inhibitors — no. (%)	15 (9.2)	15 (9.5)	30 (9.3)
Exposure to ARBs — no. (%)	11 (6.7)	11 (7.0)	22 (6.9)

Vasopressin use during 6 hr before randomization — no. (%)	113 (69.3)	111 (70.3)	224 (69.8)
Vasopressor dose — µg/kg/min¶			
Median (IQR)	0.33 (0.23–0.56)	0.34 (0.23–0.56)	0.34 (0.23–0.56)
Distribution — no. (%)			
<0.35	83 (50.9)	83 (52.5)	166 (51.7)
≥0.35 to <0.50	34 (20.9)	27 (17.1)	61 (19.0)
≥0.50	46 (28.2)	48 (30.4)	94 (29.3)

Studie ATHOS-3: výsledky

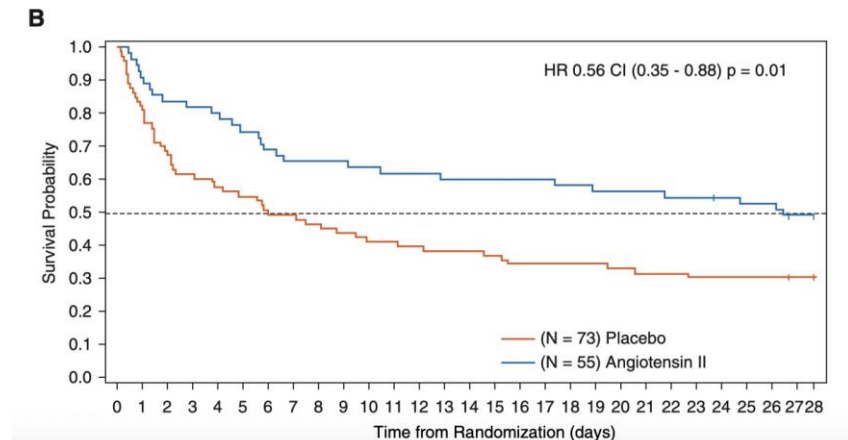
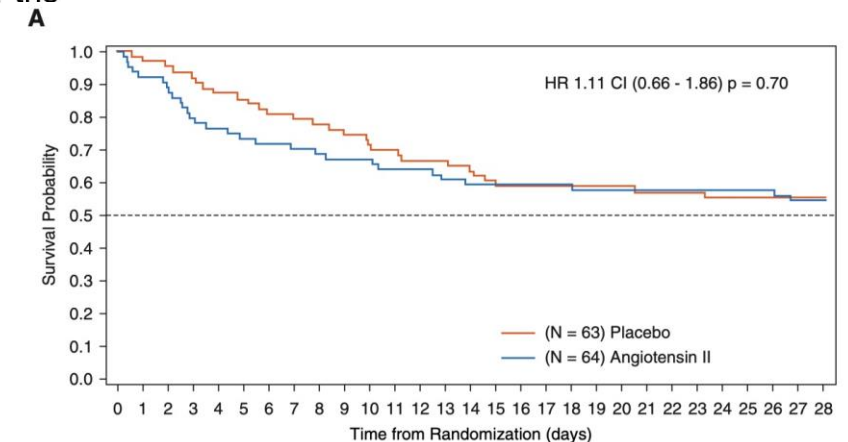
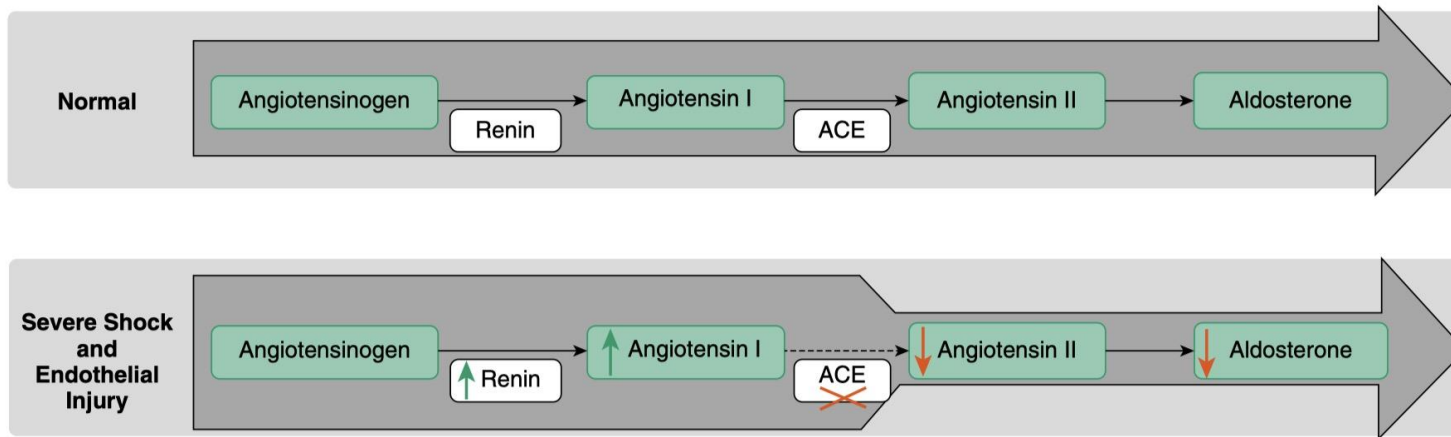
End Point	Angiotensin II (N=163)	Placebo (N=158)	Odds or Hazard Ratio (95% CI)	P Value
Primary efficacy end point: MAP response at hour 3 — no. (%)†	114 (69.9)	37 (23.4)	Odds ratio, 7.95 (4.76–13.3)	<0.001
Secondary efficacy end points				
Mean change in cardiovascular SOFA score at hour 48‡	−1.75±1.77	−1.28±1.65		0.01
Mean change in total SOFA score at hour 48§	1.05±5.50	1.04±5.34		0.49
Additional end points				
Mean change in norepinephrine-equivalent dose from baseline to hour 3¶	−0.03±0.10	0.03±0.23		<0.001
All-cause mortality at day 7 — no. (%)	47 (29)	55 (35)	Hazard ratio, 0.78 (0.53–1.16)	0.22
All-cause mortality at day 28 — no. (%)	75 (46)	85 (54)	Hazard ratio, 0.78 (0.57–1.07)	0.12



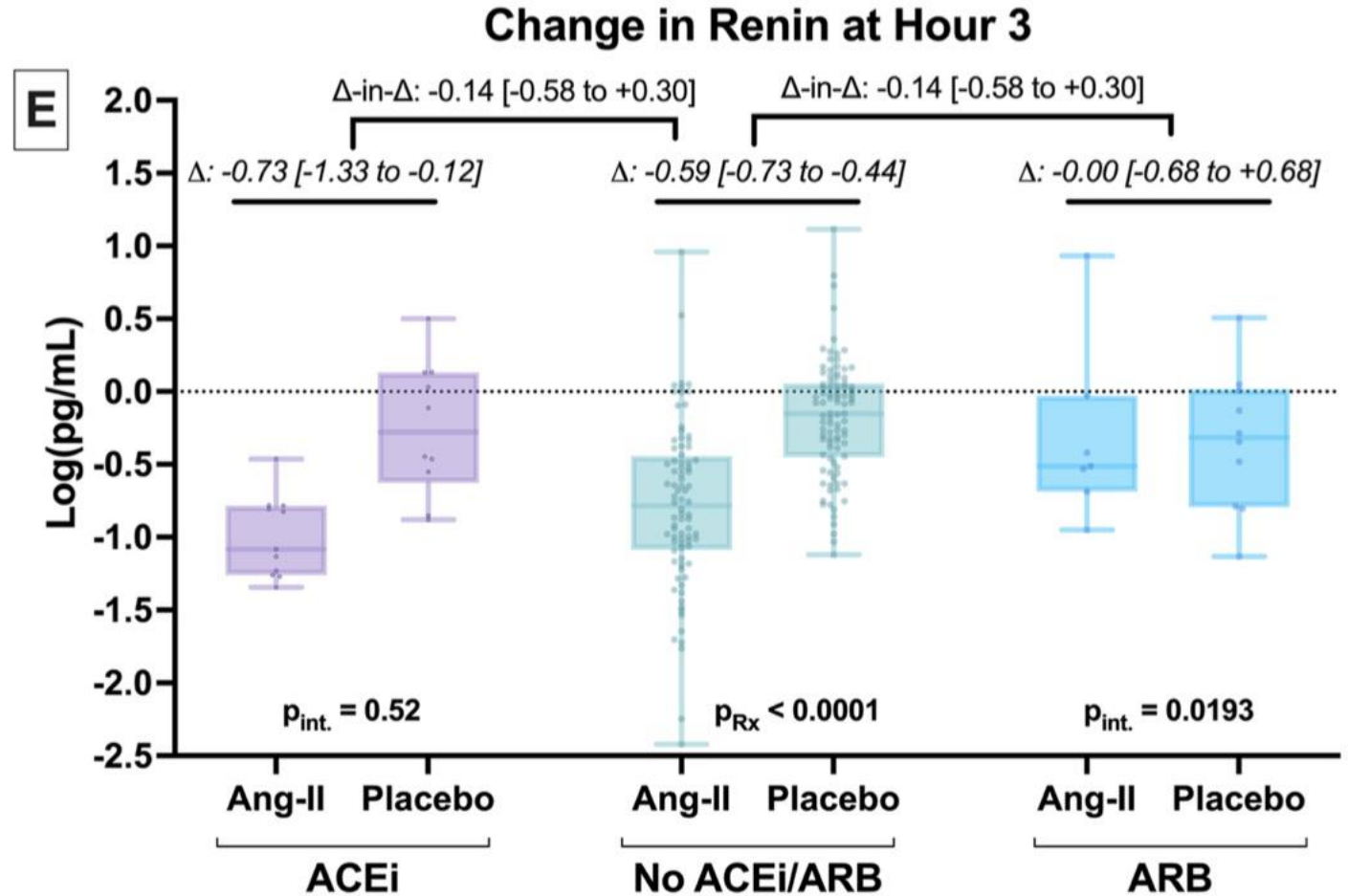
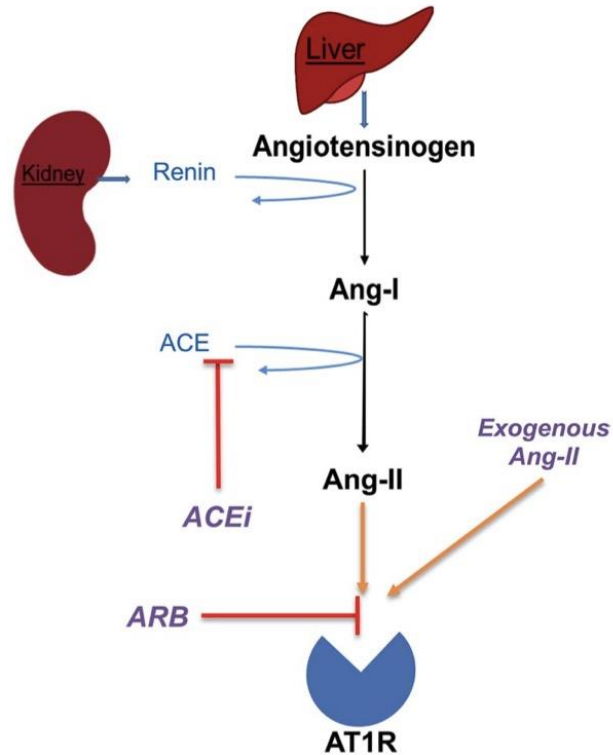
Studie ATHOS-3: post-hoc analýzy

Renin and Survival in Patients Given Angiotensin II for Catecholamine-Resistant Vasodilatory Shock A Clinical Trial

Rinaldo Bellomo^{1,2}, Lui G. Forni^{3,4}, Laurence W. Busse⁵, Michael T. McCurdy⁶, Kealy R. Ham⁷, David W. Boldt⁸, Johanna Hästbacka⁹, Ashish K. Khanna^{10,11}, Timothy E. Albertson¹², James Tumlin¹³, Kristine Storey¹⁴, Damian Handisides¹⁴, George F. Tidmarsh^{14,15}, Lakhmir S. Chawla^{14,16}, and Marlies Ostermann¹⁷; on behalf of the ATHOS-3 Investigators



Studie ATHOS-3: post-hoc analýzy



Studie ATHOS-3: post-hoc analýzy

Angiotensin-II Initial MAP Response Index of Treatment Effect (AIMRITE)

$$\text{AIMRITE} = \frac{\text{MAP}_{\text{hr1}} - \text{MAP}_{\text{hr0}}}{\text{Ang-II Dose}_{\text{hr1}}}$$

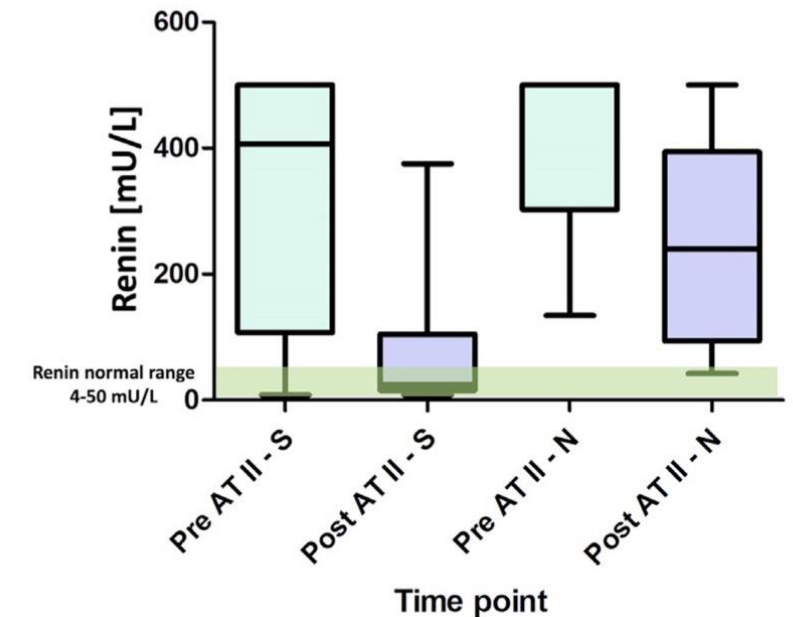
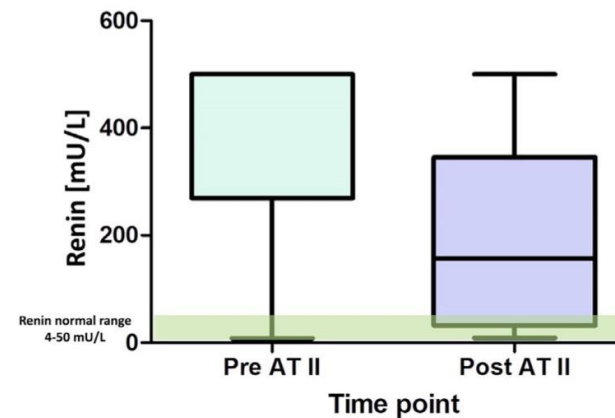
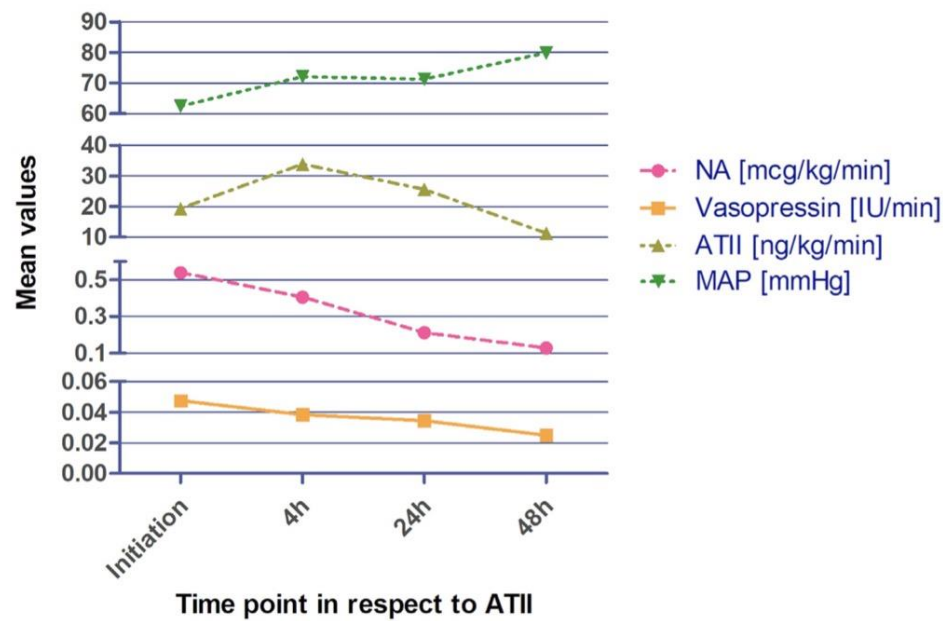
AIMRITE $\geq$ 0.90 = Responsive

1. Measure baseline MAP, immediately prior to Ang-II initiation
2. Start Ang-II infusion and titrate to MAP $\geq$ 75mmHg or 10mmHg above baseline
3. Maintain all other vasopressor infusions at constant dose
4. After 1 hour, measure MAP and Ang-II dose
5. Calculate and interpret AIMRITE

		Ang-II Responsive		Ang-II Resistant	
		PATIENT 1	PATIENT 2	PATIENT 3	PATIENT 4
Hour 0	<u>Measure</u> MAP	65 mmHg	60 mmHg	65 mmHg	60 mmHg
		↓	↓	↓	↓
Hour 1	<u>Measure</u> MAP	74 mmHg	80 mmHg	74 mmHg	80 mmHg
	Ang-II Dose	2.5 ng/kg/min	20 ng/kg/min	20 ng/kg/min	80 ng/kg/min
	<u>Calculate</u> AIMRITE	$= \frac{74 - 65 \text{ mmHg}}{2.5 \text{ ng/kg/min}} = 3.60$	$= \frac{80 - 60 \text{ mmHg}}{20 \text{ ng/kg/min}} = 1.00$	$= \frac{74 - 65 \text{ mmHg}}{20 \text{ ng/kg/min}} = 0.45$	$= \frac{80 - 60 \text{ mmHg}}{80 \text{ ng/kg/min}} = 0.25$
		Highly Sensitive to Ang-II		Minimally Responsive to Ang-II	

Intenzivní medicína: reálné zkušenosti s AT II

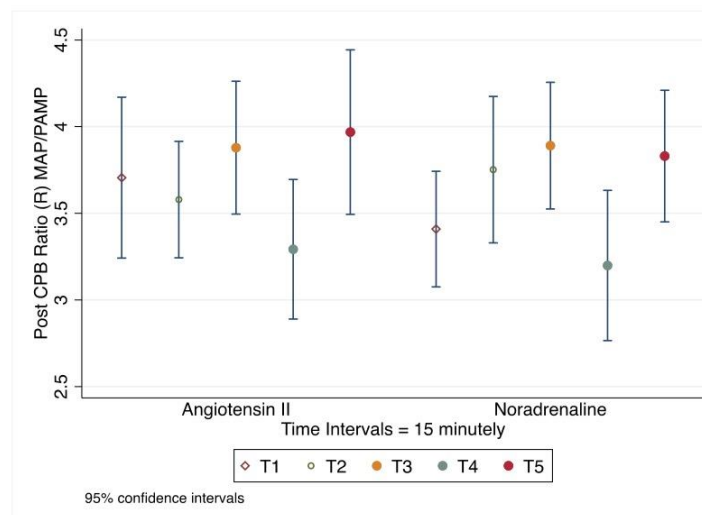
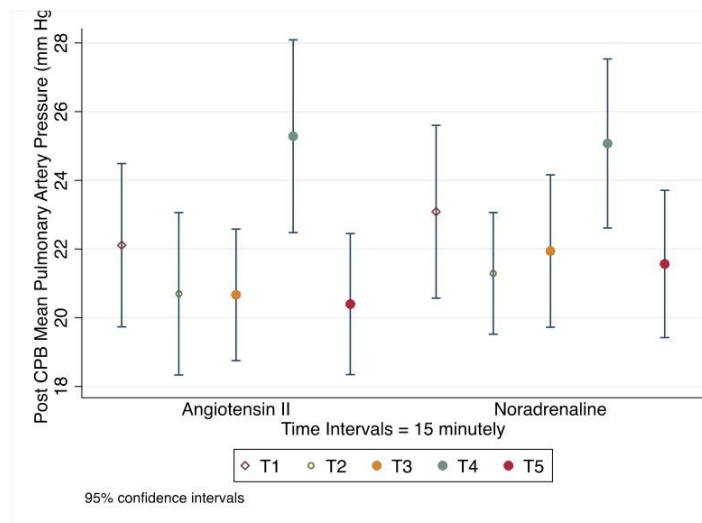
- 42 pacientů na ICU
- ✓ Distributivní šok + vazopresorická podpora NAD ≥ 0.3 ug/kg/min + VAS



Vliv AT II na plicní cévní rezistenci

■ Post-hoc analýza

- ✓ 58 pacientů
- ✓ KCH výkon s CPB
- ✓ Vyšší riziko rozvoje AKI



Original Article

The Effects of Angiotensin II versus Norepinephrine on Pulmonary Vascular Resistance in Cardiac Surgery: Post Hoc Analysis of a Randomized Controlled Trial

Jolene Lim, MBBS, MSc, FANZCA^{*,†}, Kathy Zhang, MD[‡],
Lachlan Miles, PhD, FANZCA^{†,§},
Rinaldo Bellomo, MD, PhD^{§,||,¶,***},
Tim G. Coulson, PhD, FANZCA^{*,§,1}

Author, year	No. of patients	Population	Follow-up
<i>Randomized controlled trials</i>			
Bennett SR, 2001 ⁷⁴	20	Cardiac surgery	48 hours
Chawla LS, 2014 ⁶⁶	20	Septic shock	30 days
Khanna A, 2017 ⁴	321	Mixed vasodilatory shock	28 days
Coulson TG, 2022 ⁵	60	Cardiac surgery	Hospital discharge
Sadjadi M, 2024 ⁶	63	Cardiac surgery	90 days
<i>Observational studies</i>			
Singh S, 1966 ⁷²	50	Septic shock	Hospital discharge
Leisman DE, 2020 ²²	29	COVID-19	7 days
Wong A, 2020 ^{23 a}	16	Mixed vasodilatory shock	30 days
Wieruszewski PM, 2021 ^{24 ab}	270	Mixed vasodilatory shock	30 days
Andrews L, 2022 ⁷	20	Kidney transplant	30 days
Meersch M, 2022 ²⁵	40	Cardiac surgery	28 days
Quan M, 2022 ²⁶	147	Septic shock	Hospital discharge
Serpa Neto A, 2022 ⁸	132	COVID-19	Hospital discharge
Smith SE, 2022 ^{27 ab}	162	Mixed vasodilatory shock	28 days
Ten Lohuis CC, 2022 ^{28 ab}	78	Cardiac surgery	30 days
Wieruszewski PM, 2023 ^{29 ab}	50	Mixed vasodilatory shock	28 days
Wieruszewski PM, 2023 ^{30 ab}	254	Mixed vasodilatory shock	Hospital discharge
Smith LM, 2023 ^{31 ab}	813	Mixed vasodilatory shock	90 days
See EJ , 2023 ³²	120	Mixed vasodilatory shock	90 days
Blankenship CR, 2024 ³³	69	Septic shock	28 days
<i>Case reports and case series</i>			
Cohn JN, 1965 ⁶⁷	28	Septic shock	1 hour after vasopressor discontinuation
Belle MS, 1965 ⁶⁸	1	Cardiogenic shock	11 days
Jackson T, 1993 ⁶⁹	1	Drug overdose	20 hours
Trilli LE, 1994 ⁷⁰	1	Drug overdose	10 days
Newby DE, 1995 ⁷¹	1	Drug overdose	Hospital discharge
Yunge M, 2000 ⁷³	2	Septic shock	24 days
Ahmed M, 2018 ³⁴	1	Multiple trauma	Hospital discharge
Chow JH, 2018 ^{35 ab}	1	Septic shock	41 days
Bailey DM, 2019 ³⁶	2	Septic shock	10 days
Evans A, 2019 ^{37 a}	1	Cardiac surgery	12 days
Wieruszewski PM, 2019 ^{38 ab}	4	Cardiac surgery	29 days
Wieruszewski PM, 2019 ³⁹	1	Heart and liver transplant	Surgery completion
Bobeck KA, 2020 ⁴⁰	1	COVID-19	15 days
Bui AD, 2020 ⁴¹	1	Septic shock	7 days
Chatterjee S, 2020 ⁴²	1	Thoracoabdominal aortic surgery	18 days
Coleman PJ, 2020 ⁴³	1	Septic shock	9 days
Cutler NS, 2020 ^{44 a}	1	Cardiac surgery	18 days
Ferdowsali J, 2020 ⁴⁵	1	Drug overdose	9 days
Heinicke U, 2020 ⁴⁶	6	COVID-19	16 days
Hylton DJ, 2020 ⁴⁷	1	Pheochromocytoma resection	8 days
Morselli F, 2020 ⁴⁸	7	COVID-19	28 days
Trethowan B, 2020 ^{49 ab}	1	Cardiac surgery	11 days
Wang H, 2020 ⁵⁰	1	COVID-19	9 days
Wieruszewski PM, 2020 ⁵¹	1	Mixed vasodilatory shock	5 days
Chandra R, 2021 ⁵²	1	Septic shock	Hospital discharge
Chow JH, 2021 ^{53 ab}	2	Septic shock	37 days
Cutler NS, 2021 ^{54 b}	4	Heart transplant	29 days
Ofosu-Barko K, 2021 ⁵⁵	10	COVID-19	Hospital discharge
Running K, 2021 ^{56 b}	1	Liver transplantation	14 days
Smith SE, 2021 ⁵⁷	2	Drowning	7 days
Sovic W, 2021 ^{58 a}	2	Cardiac surgery	30 days
Gutierrez GC, 2022 ⁶⁴	1	Drug overdose	9 days
Konkol SB, 2022 ⁵⁹	6	Cardiac surgery	Hospital discharge
Mohamed A, 2022 ^{60 a}	14	Cardiogenic shock	Hospital discharge
Bird S, 2023 ^{61 a}	19	Cardiac surgery	30 days
Johnson AJ, 2023 ⁷⁵	30	Cardiac surgery	28 days
Šribar A, 2023 ⁶²	1	Cardiac surgery	46 days
Tezel O, 2023 ⁶³	23	Mixed vasodilatory shock	7 days
Razdan S, 2024 ⁶⁵	1	Bilateral renal agenesis	6 months

Author, year	Maximal dose	Duration
<i>Randomized controlled trials</i>		
Bennett SR, 2000 ⁷⁴	Not reported	24 hours
Chawla LS, 2014 ⁶⁶	40 ng/kg/min	7 days
Khanna A, 2017 ⁴	40 ng/kg/min	48 hours
Coulson TG, 2022 ⁵	27.2 g/min	217.5 minutes (surgery), 5 hours (ICU)
Sadjadi M, 2024 ⁶	80 ng/kg/min	12 hours
<i>Observational studies</i>		
Singh S, 1966 ⁷²	12 g/min	105 hours
Wong A, 2020 ²³	80 ng/kg/min	15.2 hours
Wieruszewski PM, 2021 ²⁴	52 ng/kg/min	23 hours
Meersch M, 2022 ²⁵	80 ng/kg/min	8 hours
Quan M, 2022 ²⁶	80 ng/kg/min	Not reported
Serpa Neto A, 2022 ⁸	32 ng/kg/min	5 days
Smith SE, 2022 ²⁷	40 ng/kg/min	25 hours
Ten Lohuis CC, 2022 ²⁸	40 ng/kg/min	Not reported
Wieruszewski PM, 2023 ²⁹	Not reported	12 hours
Wieruszewski PM, 2023 ³⁰	40 ng/kg/min	47 hours
Smith LM, 2023 ³¹	Not reported	17 hours
See EJ , 2023 ³²	40 ng/kg/min	43 hours
<i>Case reports or case series</i>		
Belle MS, 1965 ⁶⁸	250 mg/day	7 days
Cohn JN, 1965 ⁶⁷	60 g/min	Not reported
Jackson T, 1993 ⁶⁹	18 g/min	4 hours and 50 minutes
Trilli LE, 1994 ⁷⁰	9.0 g/min	120 hours
Newby DE, 1995 ⁷¹	100 g/h	30 hours
Yunge M, 2000 ⁷³	800 ng/kg/min	7 days
Ahmed M, 2018 ³⁴	15 ng/kg/min	51 hours
Chow JH, 2018 ³⁵	80 ng/kg/min	36.5 hours
Bailey DM, 2019 ³⁶	39.1 ng/kg/min	80 hours
Evans A, 2019 ³⁷	40 ng/kg/min	13 hours
Wieruszewski PM, 2019 ³⁹	50 ng/kg/min	20 hours
Bobeck KA, 2020 ⁴⁰	Not reported	8 days
Bui AD, 2020 ⁴¹	20 ng/kg/min	6 days
Chatterjee S, 2020 ⁴²	80 ng/kg/min	40 hours
Coleman PJ, 2020 ⁴³	40 ng/kg/min	4 days
Cutler NS, 2020 ⁴⁴	15 ng/kg/min	24 hours
Ferdowsali J, 2020 ⁴⁵	10 ng/kg/min	72 hours
Hylton DJ, 2020 ⁴⁷	38 ng/kg/min	42 hours
Trethowan B, 2020 ⁴⁹	40 ng/kg/min	17.5 hours
Wang H, 2020 ⁵⁰	10 ng/kg/min	1 day
Wieruszewski PM, 2020 ⁵¹	30 ng/kg/min	16 hours
Chandra R, 2021 ⁵²	40 ng/kg/min	Not reported
Chow JH, 2021 ⁵³	80 ng/kg/min	65 hours
Cutler NS, 2021 ⁵⁴	80 ng/kg/min	34 hours
Running K, 2021 ⁵⁶	Not reported	16 hours
Smith SE, 2021 ⁵⁷	50 ng/kg/min	14.5 hours
Sovic W, 2021 ⁵⁸	25 ng/kg/min	42 hours
Gutierrez GC, 2022 ⁶⁴	80 ng/kg/min	48 hours
Konkol SB, 2022 ⁵⁹	80 ng/kg/min	20 hours
Bird S, 2023 ⁶¹	80 ng/kg/min	58.5 hours
Johnson AJ, 2023 ⁷⁵	80 ng/kg/min	9.4 hours
Šribar A, 2023 ⁶²	Not reported	24 hours
Tezel O, 2023 ⁶³	80 ng/kg/min	27 hours
Razdan S, 2024 ⁶⁵	40 ng/kg/min	10 days

Original Article

The Efficacy and Safety of Angiotensin II for Treatment of Vasoplegia in Critically Ill Patients: A Systematic Review

Yuki Kotani, MD^{*}, Martina Lezzi, MD[†], Carlotta Pia Murru, MD[†],
Ashish K Khanna, MD, FCCP, FCCM^{†,§,||},
Alexander Zarbock, MD, PhD[¶],
Rinaldo Bellomo, MD, PhD, FRACP, FCICM^{**††‡‡},
Giovanni Landoni, MD^{†,§§,¹}

Conclusions: Intravenous angiotensin II has been reported in almost 3000 critically ill patients with diverse types of shock. Despite unclear mortality impacts, angiotensin II seems to confer beneficial effects on several organ systems and RAS derangements, without increasing adverse events.

Transplantace srdce

Case Summary

	Case 1	Case 2	Case 3	Case 4
Age (y), sex	62, male	61, male	69, male	60, male
Primary diagnosis	NICM	ICM	ICM	NICM
Prior treatment	HM3 LVAD BTT	CABG, HM3 LVAD BTT	CABG twice, long-term milrinone	HM3 LVAD BTT
Sternotomy #	2	3	3	3
Ischemia time	259 min	181 min	194 min	226 min
Just prior to starting ANG-2:				
Vasopressor support	NE 0.29 µg/kg/min AVP 0.06 units/min	NE 0.11 µg/kg/min AVP 0.04 units/min EPI 0.03 µg/kg/min	NE 0.29 µg/kg/min AVP 0.03 units/min EPI 0.06 µg/kg/min	NE 0.06 µg/kg/min AVP 0.04 units/min EPI 0.04 µg/kg/min
MAP (mean of last 3)	59 mmHg	59 mmHg	70 mmHg	71 mmHg
ANG-2:				
Time initiated	1 h after wean from CPB	1.5 h before wean from CPB	10 hours after ICU arrival, 13 h after CPB	45 min after wean from CPB
Dose range	10-60 ng/kg/min	2.5-60 ng/kg/min	5-30 ng/kg/min	5-80 ng/kg/min
Duration	12 h	34 h	20 h	29 h
Adjunctive therapies for vasoplegia:				
Prior to ANG-2	-	-	Hydroxocobalamin 5 g	-
After ANG-2	Methylene blue, 2 mg/kg	Methylene blue, 1.5 mg/kg Hydroxocobalamin, 5 g	-	-
Other hemodynamic support required during ICU stay	IABP Milrinone Dobutamine Epinephrine Inhaled epoprostenol	IABP Milrinone Inhaled epoprostenol	IABP Milrinone Inhaled epoprostenol	Milrinone Inhaled epoprostenol
Major complications	Open chest owing to poor hemodynamics Ischemic optic neuropathy Subdural hematoma IABP insertion site pseudoaneurysm	Re-exploration (bleeding) Open chest owing to poor hemodynamics Renal failure Respiratory failure, reintubation VAP Femoral cannulation site dehiscence	Liver injury	Agitated delirium
Discharge date	POD 28	POD 49	POD 15	POD 22



Case Report Angiotensin II for Critically Ill Patients With Shock After Heart Transplant

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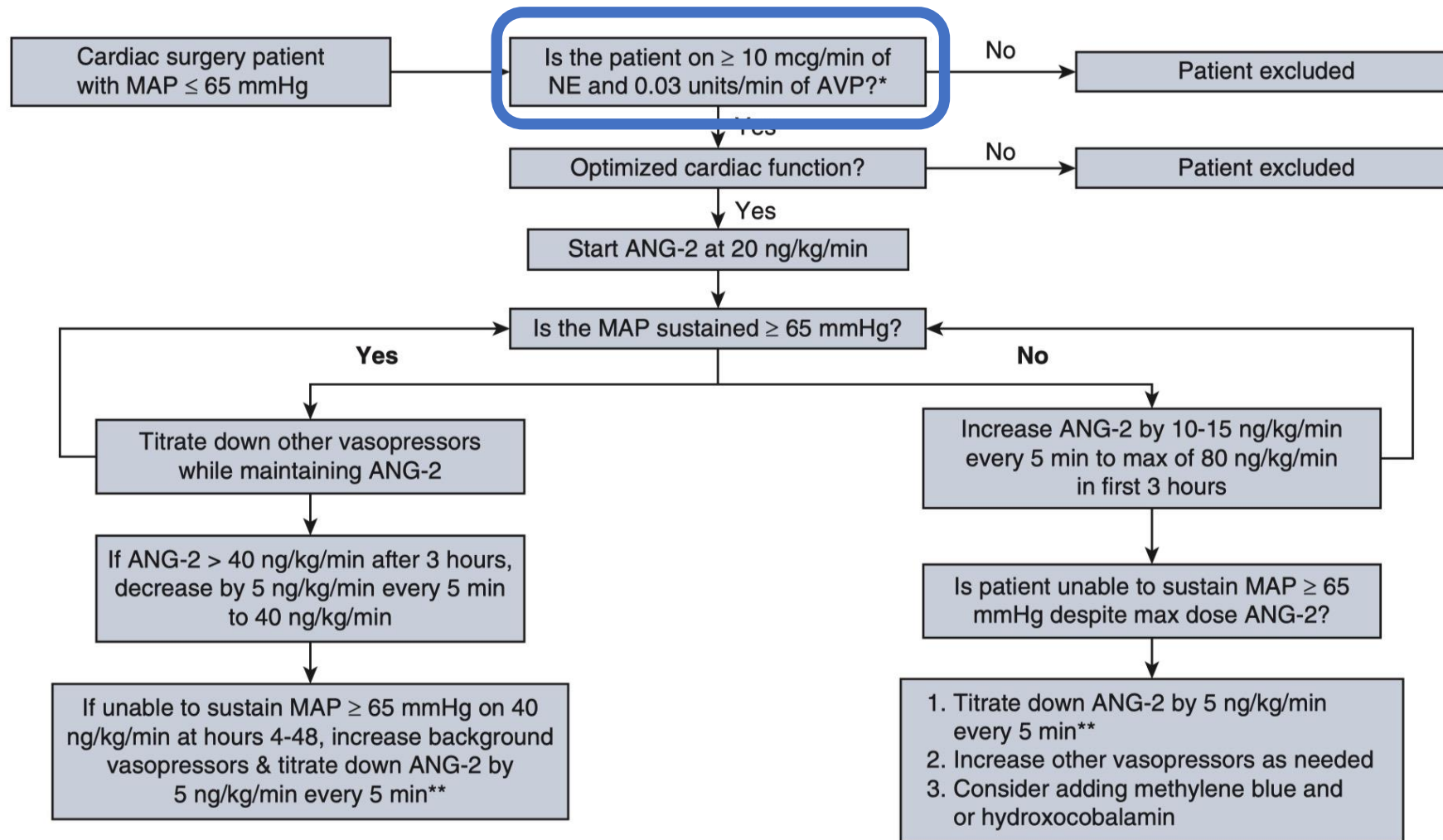
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Protokoly pro kardiochirurgické pacienty



Protokoly pro kardiochirurgické pacienty

Proposal for the use of angiotensin II in distributive shock after extracorporeal circulation – position paper of the Section of Intensive Care Medicine and the Section of Cardiothoracic Anaesthesiology of the Polish Society of Anaesthesiology and Intensive Therapy



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Kardiochirurgia i Torakochirurgia Polska 2024; 21 (2): 96-98

Haemodynamic criteria of inclusion:
 $\text{MAP} < 65 \text{ mm Hg} + \text{SVR} < 500 \text{ dyn} + \text{NED} \geq 0.2 \text{ (0.3)} \mu\text{g/kg/min}$
Haemodynamic criteria of exclusion: $\text{CI} < 2.2 \text{ l/min/m}^2$
or $\text{mPAP} > 45 \text{ mm Hg}$

Solution preparation: 2.5 mg AT in 500 ml 0.9% NaCl (5,000 ng/ml)

Starting AT dose: (10) 20 ng/kg/min + escalation to reach
MAP 75 mm Hg

Maximum AT dose: 80 ng/kg/min in 3 h

Maintenance AT dose: $\leq 40 \text{ ng/kg/min}$ up to 48 h to reach MAP
+75 mm Hg + reduction of vasoactive drugs until
NED $0.1 \mu\text{g/kg/min}$

AT de-escalation: 5 ng/kg/min every 5 min if MAP $> 65 \text{ mm Hg}$

Doporučené postupy

V. Indikace a podávání angiotensinu II

6. Nitrožilní podání ang-II průkazně snižuje potřebu katecholaminů u pacientů s vazoplegickým šokem nereagujících na standardní léčebné postupy (zahrnuje obvykle podávání noradrenalinu v kombinaci s vazopresinem).
7. Nitrožilní podání ang-II by mělo být zváženo u pacientů
 - a) s klinickým fenotypem vazoplegie (hypotenze, nízká systémová vaskulární rezistence, normální nebo zvýšený srdeční výdej),
 - b) nereagujících na standardní vazopresory a
 - c) kde jsou vyloučeny všechny ostatní odstranitelné příčiny s možným podílem na rozvoji vazoplegie.
8. Současný stav odborného poznání neumožňuje formulovat jednoznačné stanovisko k měření hladin reninu v procesu stanovení indikace podávání ang-II.
9. Před zahájením podávání ang-II by mělo být vždy vyhodnoceno aktuální individuální riziko hluboké žilní trombozy a zvážen poměr přínosu a rizika zahájení profylaxe tromboembolické choroby.

Angiotensin II – mechanismus účinku, současná evidence a stanovisko mezioborové pracovní skupiny k jeho použití na pracovištích intenzivní péče

Balík M.¹, Beneš J.², Černý V.³⁻⁸, Kula R.⁹, Matějovič M.¹⁰, Říha H.¹¹, Tencer T.¹²

MAP < 65 mmHg
NAD ≥ 0,5 µg/kg.min
AVP ≥ 0,04 IU/kg.h
Zvýšená hladina reninu
Léčba ACEI
VA-ECMO
ECC



Iniciální dávka:
2–20 ng/kg.min, titrace
v 5min intervalech do
max. 80 ng/kg.min po
první 3 h



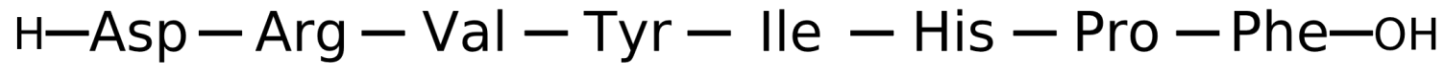
Udržovací dávka od 3 h:
1,25–40 ng/kg.min

Naše zkušenosti

- 24 pacientů (10/2023 - 09/2025)
- Všechny typy KCH výkonů
 - ✓ V-A ECMO 8x
 - ✓ HM3 LVAD 4x
 - ✓ OTS 5x
- RES (vs. operační sál [během CPB])
- Zahájení infuze: dávka ostatních vazopresorů (NAD + VAS)
- Délka trvání infuze: 12-48 h (72-96 h)
- Dávka po iniciální stabilizaci hemodynamiky: 10-30 ng/kg/min
- Účinek

Angiotensin II

- ✓ Poločas AT II 15 s
- ✓ Poločas Giapreza® <1 min



- Komu v kardiochirurgii podávat AT II?
- Kdy zahájit aplikaci AT II (dávka noradrenalinu/vazopresinu)?
- Vazopresor 1./2./3. linie (multimodální strategie)?
- Dostupnost rychlého stanovení plazmatické koncentrace reninu?

