



ECT – anesthesia:

1. Typical anesthetic drugs (p, e, m ,t)

a. Ketofol - two stones to catch one bird ?

2. Some typical problems / solutions ...

- **ASTI** a.
- Oxygen b.
- PAS / PIA C.
- Cardiac d.

substance	typical dose range (mg/kg)	anticonvulsive effect (relative)	remarks	26
methohexital	0,75-1.0	1-2	former gold standard, cardiovascular depression	Black Box Warn *
thiopental	2-5	2	cardiovascular depression	Rote-Hand-Briefe
propofol	1-2	3	shorter seizures, higher seizure threshold	1
etomidate	0.2-0.3	0	myocloni	***
S-ketamine	0.5-1.5	0	low doses pro-psychotic, higher blood pressure	de la
alfentanil	0.01-0.015	0	longer time of apnoe, cardiovascular depression	-
remifentanil	0.001-0.008	1	similar to alfentanil ?	mark

Hikma Pharmaceuticals stopped the production of methohexital 2019

Good Manufacturing Practice (GMP) problems at Lampugnani Pharmaceutici SPA

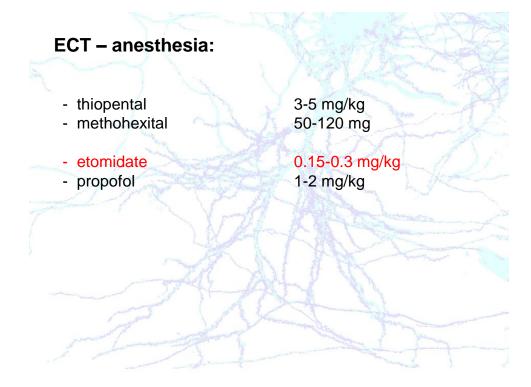
In Germany: more or less obsolete for critically ill patients and for non-single induction use

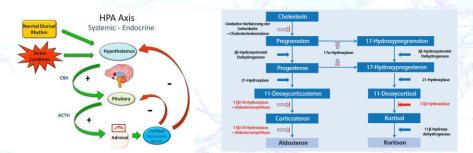
Adapted from:

Folkerts HW.

Electroconvulsive therapy. Indications, procedure and treatment results Nervenarzt. 2011 Jan;82(1):93-102

Swartz CM Electroconvulsive and neuromodulation therapies. 2009 Cambridge Univ, Cambridge New York Melbourne





Etomidate: to use or not to use for endotracheal intubation in the critically ill? Smischney NJ, Kashyap R, Gajic O. J Thorac Dis. 2015 Sep;7(9):E347-9.

Etomidate for intravenous induction of anaesthesia Dumps C, Bolkenius D, Halbeck E. Anaesthesist. 2017 Dec;66(12):969-980.

Optimistic viewpoint:

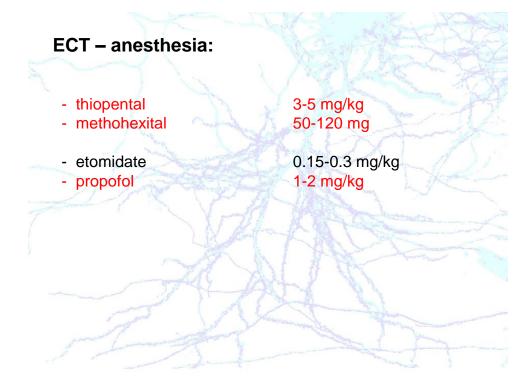
Raeder J., Curr Opin Anaesthesiol. 2019 Sep 9. [Epub ahead of print] Procedural sedation in ambulatory anaesthesia: what's new?

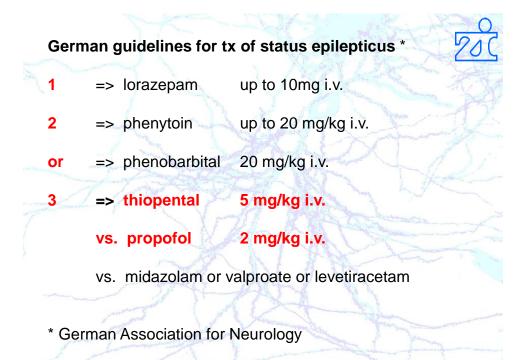
This debate is ongoing ...

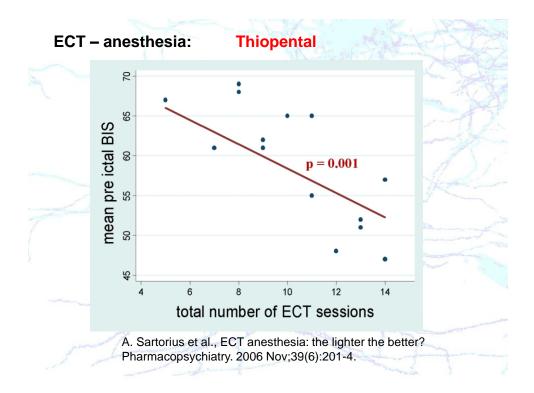
... and of course there are no long term studies in ECT patients...

The theoretical problem of a cumulative risk remains, because of a "chronic" HPA suppression with repeated use of etomidate.

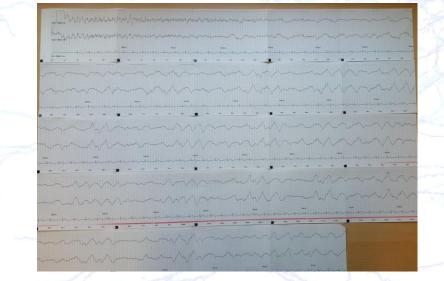
At least in Germany anesthesiologists become more and more "careful" with the use of etomidate in ECT.



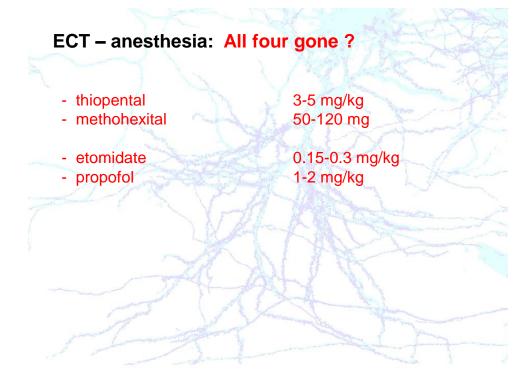




Courtesy of Michael Guhra, Bielefeld



32 years, "always had good seizures", now 250 mg propofol



ECT – anesthesia: All four gone ?

- thiopental
- methohexital
- etomidate
- propofol

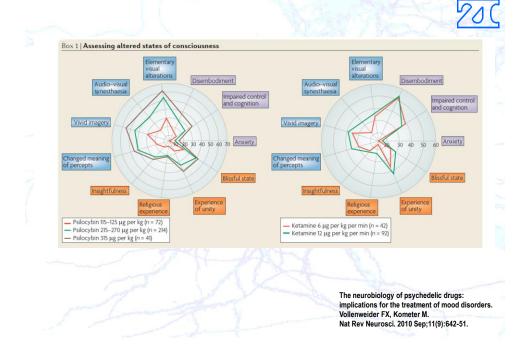
- 3-5 mg/kg 50-120 mg
- 0.15-0.3 mg/kg 1-2 mg/kg

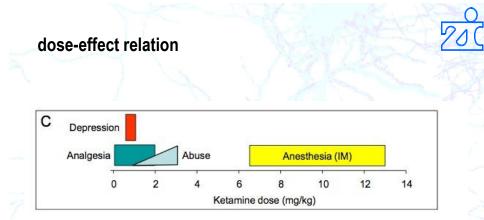
Of course not, but all have major drawbacks.

What about ketamine ?

ketamine in general anesthesia:

- is listed as an essential drug by the WHO
- often used in emergeny medicine
- treatment of status asthmaticus
- analgesia of intubated patients
- preferred for childs and ado's
- still in use for general and regional anesthesia - alone and in combination with hypnotics
- off-label for chronic pain patients





Glue et al., Biol Psychiatry, 2011

ketamine racemate in mg/kg bw i.m. as bolus

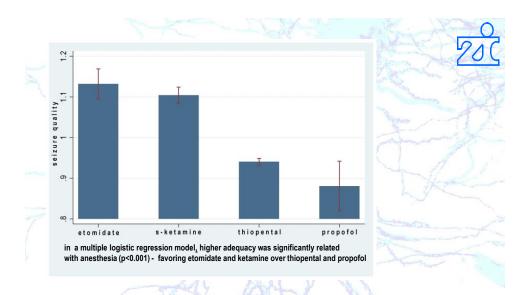
ECT and ketamine

pros:

- 1. Ketamine probably posseses a unique intrinsic antidepressive potential
- 2. Ketamine has no anticonvulsive action
- 3. Ketamine may posses neuroprotective properties as an NMDA-antagonist

cons:

- 1. Ketamine acts non-depressively on the cardio-vascular systeme (like e.g. barbiturates)
- 2. Ketamine dose-dependently induces psychiatric side-effects (basically derealisation and depersonalisation, which can lead to anxiety)



Impact of ketamine, etomidate, thiopental and propofol as anesthetic on seizure parameters and seizure quality in electroconvulsive therapy: A retrospective study

Carolin Hoyer, Laura Kranaster, Christoph Janke, Alexander Sartorius Eur Arch Psychiatry Clin Neurosci 2014 Apr;264(3):255-61.

		Ket			Con			Std. Mean Difference	Std. Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
Abdallah 2012	22.9	12.4	7	20.4	11.1	7	3.4%	0.20 [-0.85, 1.25]	
Nizadeh 2015	16.27	6.4	22	14.77	6.82	20	6.7%	0.22 [-0.38, 0.83]	
nderson 2017	17.2	11.6	27	15	10.4	32	7.8%	0.20 [-0.32, 0.71]	
hen 2017	10.07	5.96	63	12.06	5.81	64	10.1%	-0.34 [-0.69, 0.01]	
ernie 2017	13.5	9.32	16	8.41	4.7	17	5.7%	0.68 [-0.03, 1.38]	
larventausta 2013	10	10.93	16	10.2	9.05	16	5.8%	-0.02 [-0.71, 0.67]	
(uscu 2015	4.5	2.58	38	3.7	1.6	20	7.4%	0.34 [-0.20, 0.89]	
.00 2012	14.28	10.34	22	16.78	10.49	24	7.0%	-0.24 [-0.82, 0.34]	
Rasmussen 2014	22.08	8.11	21	24.45	7.7	17	6.3%	-0.29 [-0.94, 0.35]	
Rybakowski 2016	12.5	6.05	30	15.9	6.6	15	6.5%	-0.54 [-1.17, 0.09]	
alehi 2015	8.32	5.17	80	10.53	7.87	80	10.6%	-0.33 [-0.64, -0.02]	
'oosefi 2014	17.2	2.46	17	17.71	2.46	14	5.7%	-0.20 [-0.91, 0.51]	
hang 2017.	15.55	7	43	17.77	6.47	34	8.6%	-0.32 [-0.78, 0.13]	
(hong 2016	6.55	1.34	60	8.2	1.9	30	8.4%	-1.06 [-1.52, -0.59]	
otal (95% CI)			462			390	100.0%	-0.17 [-0.39, 0.06]	•
Heterogeneity: Tau ² =	= 0.10; C	hi² = 30	.42, df =	= 13 (P =	= 0.004)	: P= 5	7%		-2 -1 0 1 2



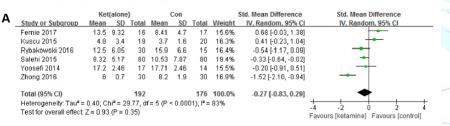


Fig. 4. (A): Meta-analysis of depressive symptoms with ketamine alone at the end of ECT course.

But:

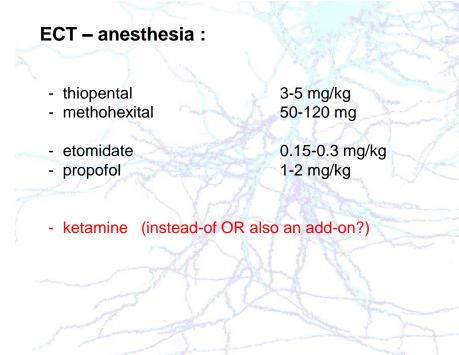
Study	Cognitive evaluation	Findings	
Shams et al., 2015	"Cognitive Performance Recovery Time" after each ECT	Ketamine group had a shorter cognitive performance recovery time compared to propofol group.	
Anderson et al., 2017	Hopkins Verbal Learning Test-Revised (HVLT-R-R)R; originally Controlled Oral Word Association Test (COWYT; Autobio spaphical Memory Interview-Short Form (AMI-SF); Medical College of Georgia Complex Figure Test (MoGCIPT), Inicial digit span forwards and backward; Self-reported Global Self Evaluation of Memory (GSE-Wy)	No significant differences between ketamine + propolol or thiopental and propolol or thiopental groups	£
Chen et al., 2017	MMSE; Wechsler Memory Scale-Chinese Revision (WMS-RC)	MMSE score was significantly lower in the ketamine group compared to the propofol group; WMS-RC score was significantly lower in propofol group compared to ketamine + propofol group	-
loo et al., 2012	Medical College of Georgia Complex Figure (CFT); Hopkins Verbal Leaming Test (HVLT); Controlled Oral Word Association Test (COWAT); symbol Digit Modallius Test (SDMT); Woodcock Johnson Cross-Out Test; Autobiographical Memory Interview—short form (AMI- SF).	No significant differences between ketamine + thiopental and thiopental groups	
Zhong et al., 2016	The Word Fluency Test; the Digit Symbol Test; the Digit Span test; the Wisconsin Card Sorting test (WCST); the Tower of Hanoi; the Trail Making Test (TMT); the Visual Regeneration Test.	Ketamine group showed a lower degree of executive cognitive impairment compared to the ketamine + propofol and propofol groups	
Zhang et al., 2017	Speed of Processing (SoP), Attention/Vigilance (AV); Working Memory (WM); Verbal Learning (VrH Lrng); Visaal Learning (VrL Hrng) Reasoning and Problem Solving (RPS) Social Cognition (SC)	No significant difference was found on the MCCB between the propool group and the ketamine plus propool group	
Femie et al., 2017	Cambridge Automated Neuropsychological Test Battery Spatial Recognition Memory Task (CANTAB SRM)	No significant difference was found on the CANTAB SRM between the propofol group and the ketamine group	
roosefi et al., 2014	Mini-Metal State Examination (MMSE)	A significantly better cognitive performance was evident in ketamine- receiving group	
Rasmussen et al., 2014	Mini-Metal State Examination (MMSE)	No significant difference was found in MMSE	
ay-Griffith et al., 2017	Mini-Metal State Examination (MMSE)	No significant difference was found in MMSE	
tybakowski et al., 2016	Tests assessing visual-spatial function Tests assessing verbal auditory function Tests assessing working memory and executive function	No difference was found in the test of visual-spatial function. Impairment of verbal memory and verbal fluency were greater with ketamine.	

Adjunctive ketamine and electroconvulsive therapy for major depressive disorder: A meta-analysis of randomized controlled trials.

Zheng W, Li XH, Zhu XM, Cai DB, Yang XH, Ungvari GS, Ng CH, Ning YP, Hu YD, He SH, Wang G, Xiang YT. J Affect Disord. 2019 May 1;250:123-131.

To conclude so far, ketamine is

- Probably not as side effectively as it was feared
- Probably not more, but definitely not less effective as the grand old four (metho, thio, propo and eto)
 - Our experience is that we need less charge for ketamine which explains no difference in response rates, but still could lead into less cognitive side effects (still has to be verified)
- no study has controlled for mean charge so far





propofol plus ketamine = ketofol:

Mind the order :

ketofol:

first propofol - followed by ketamine !

Own experiences with ketofol:

- anesthesiologists are excited



- less time in recovery room
- propofol is still critical regarding seizure threshold/induction
- But how to mix propofol and ketamine ???

ECT – anesthesia: propofol+ketamine 1:1?

Acta Neuropsychiatr. 2018 Apr;30(2):61-69.

Anaesthesia for electroconvulsive therapy - new tricks for old drugs: a systematic review.

Stripp TK, Jorgensen MB, Olsen NV.

OBJECTIVE:

The objective of this review is to investigate existing literature in order to delineate whether the use of anaesthesia and timing of seizure induction in a new and optimised way may improve the efficacy of electroconvulsive therapy (ECT). METHODS:

PubMed/MEDLINE was searched for existing literature, last search on 24 June 2015. Relevant clinical studies on human subjects involving choice of anaesthetic, ventilation and bispectral index (BIS) monitoring in the ECT setting were considered. The references of relevant studies were likewise considered.

RESULTS:

Propofol yields the shortest seizures, etomidate and ketamine the longest. Etomidate and ketamine+propofol 1 : 1 seems to yield the seizures with best quality. Seizure quality is improved when induction of ECT is delayed until the effect of the anaesthetic has waned - possibly monitored with BIS values. Manual hyperventilation with 100% O2 may increase the pO2/pCO2-ratio, which may be correlated with better seizure quality. CONCLUSION:

Etomidate or a 1 : 1 ketamine and propofol combination may be the best method to achieve general anaesthesia in the ECT setting. There is a need for large randomised prospective studies comparing the effect of methohexital, thiopental, propofol, ketamine, propofol+ketamine 1 : 1 and etomidate in the ECT treatment of major depressed patients. These studies should investigate safety and side effects, and most importantly have antidepressant efficacy and cognitive side effects as outcome measures instead of seizure quality.

ECT: a new look at an old friend

Pavan Kumar Kadiyala^a and Lakshmi Deepthi Kadiyala^b

KEY POINTS

- ECT is improving into a new form that may be perceived with a lower degree of social stigma.
- Anesthesia and augmentation strategies have a significant influence on clinical efficacy and tolerability of ECT. Etomidate, or a ketamine-propofol combination, may be the first choice. Dexmedetomidine or remifentanil may be added in selected patients.
- Hyperventilation protocols and ASTI influence the clinical outcome of ECT.
- Refinements in stimulus parameters and electrode placements leading to increased focality have led to a reduction of cognitive adverse effects. RUL brief pulse ECT represents an acceptable first-line form of ECT.
- EEG ictal indices (specifically mid-ictal amplitude, postictal suppression) during ECT procedure should be monitored for therapeutic adequacy of seizure.

propofol+ketamine 1 : 1 ?

Curr Opin Anaesthesiol. 2018 Aug;31(4):453-458.

Take home for ketofol:

- reduce both, dose of k and dose of p
- apply p and then k

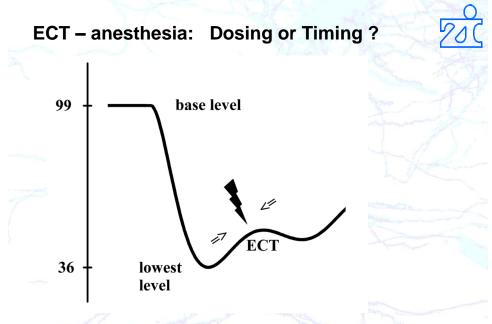
- use p : k = 1:3 (OR p : S-k = 1:1.5)

ECT – anesthesia:

- 1. Typical anesthetic drugs (p, e, m,t)
 - a. Ketofol two stones to catch one bird ?

2. Some typical problems / solutions ...

- a. ASTI (anesthesia-to-stimulation time interval)
- b. Oxygen
- c. PAS/PIA
- d. Cardiac

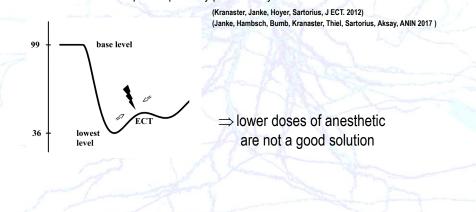


bispectrum (BIS) as a surrogate of the depth of the induced anesthesia





- dose = 0 (at unmodified ECT) results in post ictal agitation (PIA) rates of 10-50% (Andrade, Shah, Tharyan et al., Indian J Psychiatry. 2012)



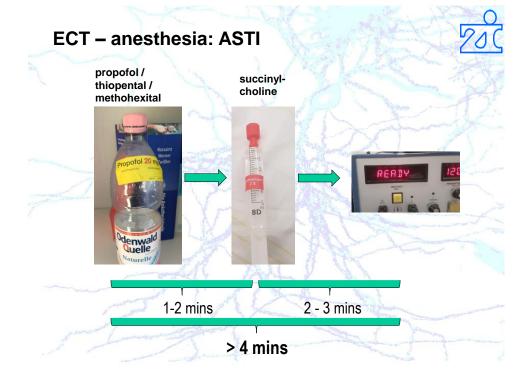
- PIA is in an individual patient perfectly predicted by BIS

ECT – anesthesia: Timing !!!

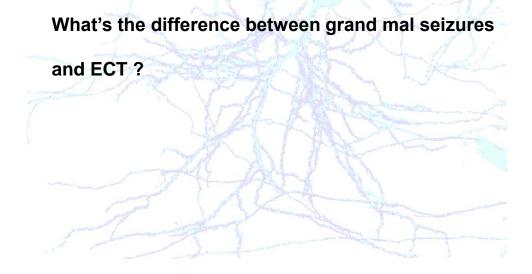
- ECT anesthesia: the lighter the better? Sartorius A, Muñoz-Canales EM, Krumm B, Krier A, Andres FJ, Bender HJ, Henn FA. Pharmacopsychiatry. 2006 Nov;39(6):201-4.

- The Anaesthetic-ECT Time Interval in Electroconvulsive Therapy Practice--Is It Time to Time? Gálvez V, Hadzi-Pavlovic D, Wark H, Harper S, Leyden J, Loo CK. Brain Stimul. 2016 Jan-Feb;9(1):72-7.

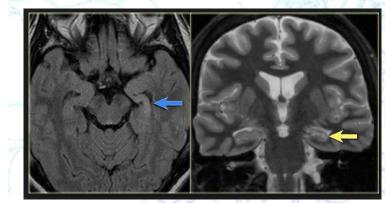
- The influence of the anesthesia-to-stimulation time interval (ASTI) on seizure quality parameters in electroconvulsive therapy. Jorgensen A, Christensen SJ, Jensen AEK, Olsen NV, Jorgensen MB. J Affect Disord. 2018 Apr 15;231:41-43.



ECT – anesthesia: Oxygen



ECT – anesthesia: Oxygen

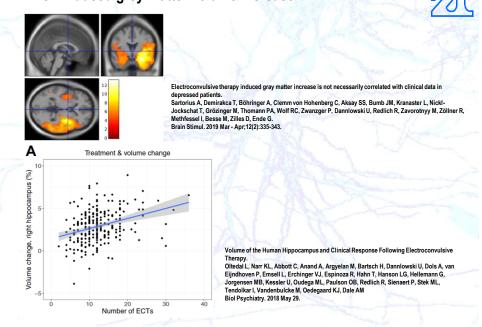


35-year-old patient with refractory temporal lobe epilepsy.

MR shows subtle hyperintensity of the left hippocampus on the axial FLAIR (blue arrow) and atrophy of the left hippocampus on coronal images (yellow arrow).



ECT induced grey matter volume increase



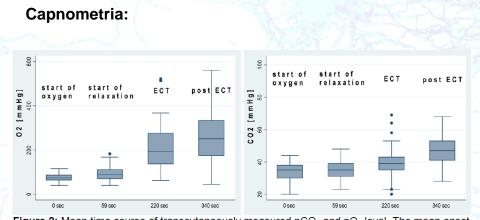
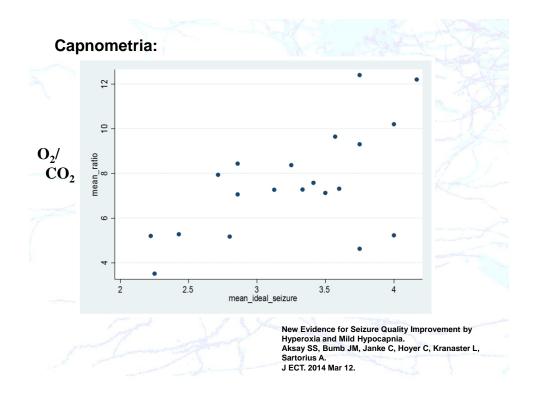


Figure 2: Mean time course of transcutaneously measured pCO₂ and pO₂ level. The mean onset of (pre-)oxygenation, muscle relaxation, start of ECT and 2 minutes post ECT are labeled.

New Evidence for Seizure Quality Improvement by Hyperoxia and Mild Hypocapnia. Aksay SS, Bumb JM, Janke C, Hoyer C, Kranaster L, Sartorius A. J ECT. 2014 Mar 12.



ECT – anesthesia: Oxygen

win –win:

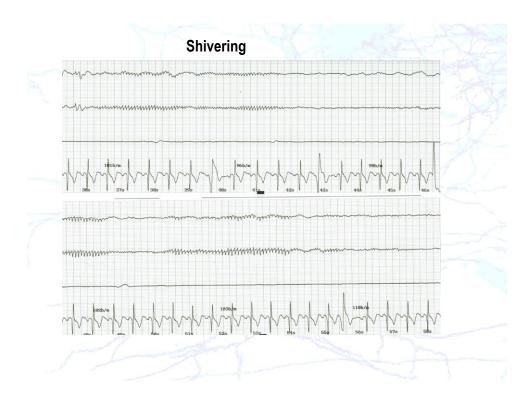
- O₂ makes the procedure safe

- O₂ lowers seizure threshold

Charles Kellner: "The green gas is the good one !"

ECT – anesthesia: PAS and PIA

Or problems with "movements" peri-ECT ...





Postanesthetic shivering (PAS) is shivering after anesthesia

is not fasciculating, is not myocloni, is not restless legs !

The intensity of PAS may be graded using the scale described by Crossley and Mahajan:

Table	1.	The	shivering	classification.
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Grade	Description
0	No shivering.
I	No visible muscle activity, but one or more of piloerection, peripheral vasoconstriction or peripheral cyanosis (other causes excluded).
2	Muscular activity in only one muscle group.
3	Moderate muscular activity in more than one muscle group, but not generalised shaking.
4	Violent muscular activity that involves the entire body.

The intensity of postoperative shivering is unrelated to axillary temperature. Crossley AW, Mahajan RP. Anaesthesia. 1994 Mar;49(3):205-7

Treatment of PAS

1. clonidine

- 2. dexmedetomidine
- 3. mivacurium instead of succinylcholine
- 4. probably more often with barbiturates / propofol and less with ketamine

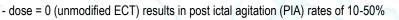
What is the place of clonidine in anesthesia? Systematic review and meta-analyses of randomized controlled trials. Sanchez Munoz MC, De Kock M, Forget P. J Clin Anesth. 2017 May;38:140-153. Review.

Systematic Quality Assessment of Published Antishivering Protocols. Choi KE, Park B, Moheet AM, Rosen A, Lahiri S, Rosengart A. Anesth Analg. 2017 May;124(5):1539-1546. Review.

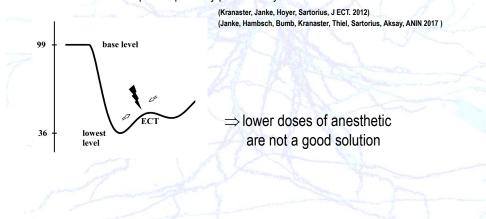
Efficiency and safety of ondansetron in preventing postanaesthesia shivering. He K, Zhao H, Zhou HC. Ann R Coll Surg Engl. 2016 Jul;98(6):358-66. Review.

Effectiveness of dexmedetomidine use in general anesthesia to prevent postoperative shivering: a systematic review. Hoffman J, Hamner C. JBI Database System Rev Implement Rep. 2016 Jan 15;13(12):287-313. Review.

ECT - anesthesia: post ictal agitation (PIA)



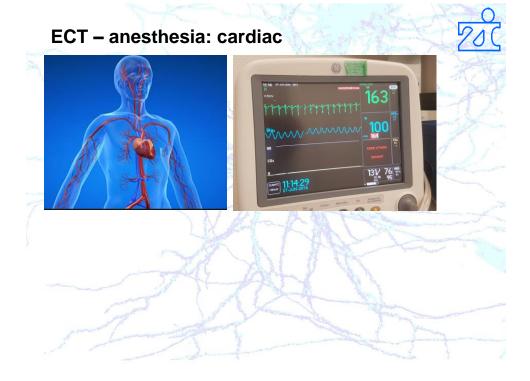
(Andrade, Shah, Tharyan et al., Indian J Psychiatry. 2012)



- PIA is in an individual patient perfectly predicted by BIS

ECT – anesthesia: post ictal agitation (PIA)

- Do not restrain !!! (=> otherwise increase of PIA)
- Keep everything calm and use as less physical limitation as possible
- Self limitating in most cases within 20 mins
- Severe forms: Escalate staff
- Severe forms: Use i.v. diazepam e.g. 10mg
- Increase dose of anesthetic next ECT



hypersalivation / sialorrhoe

- Former times:

atropine, which is basically obsolete. Why ?

- Today:

glycopyrrolate as muscarinic receptor antagonist

- Both reduce hypersalivation (parasympatholytic)
- atropine reduces initial bradycardia, but increases ictal hypertension *

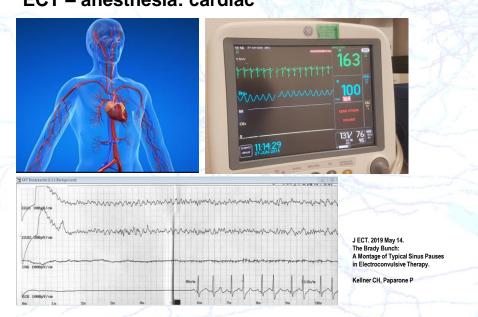
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* Psychiatry Res. 2019 Jan;271:239-246 Electro convulsive therapy: Modification of its effect on the autonomic nervous system using anti-cholinergic drugs. Christensen STJ, Staalso JM, Jørgensen A, Weikop P, Olsen NV, Jørgensen MB.

ЮΗ

H₃C

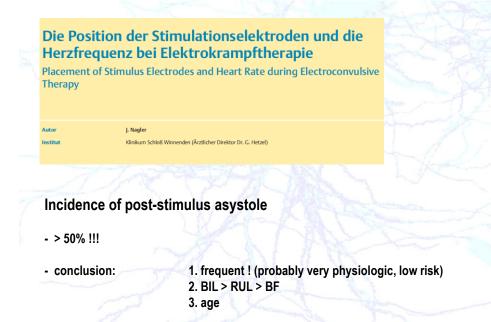
H₃C



ECT – anesthesia: cardiac



Asystolia appears shorter in our printout (printout starts at the end of charge delivery!)





	Pulsewi	Electrode placement						
Covariate	В	S.E.	р	OR	В	S.E.	p	OR
Electrode placement								
RUL vs. BF					5.334	1.333	0.000 ^b	207.239
RUL vs. BT					2.158	0.774	0.005 ^b	8.654
BT vs. BF					3.176	1.348	0.018°	23.947
Pulsewidth								
1.0 vs. 0.3	3.818	1.161	0.001	45.527				

