

# Differenzierte Opioidtherapie

Einsatz bei Komorbiditäten



Wiener  
Gesundheitsverbund



Für die  
Stadt Wien

**Beitrag: DIPIDOL<sup>®</sup> (Piritramid) Ampullen – unsachgemäßer Gebrauch**

Jede Verdünnung von Piritramid stellt in Österreich allerdings eine Off-Label-Anwendung dar.

**Wir empfehlen daher im Moment eine Umstellung auf Vendal<sup>®</sup> (Morphinhydrochlorid) 10 mg – Ampullen.**

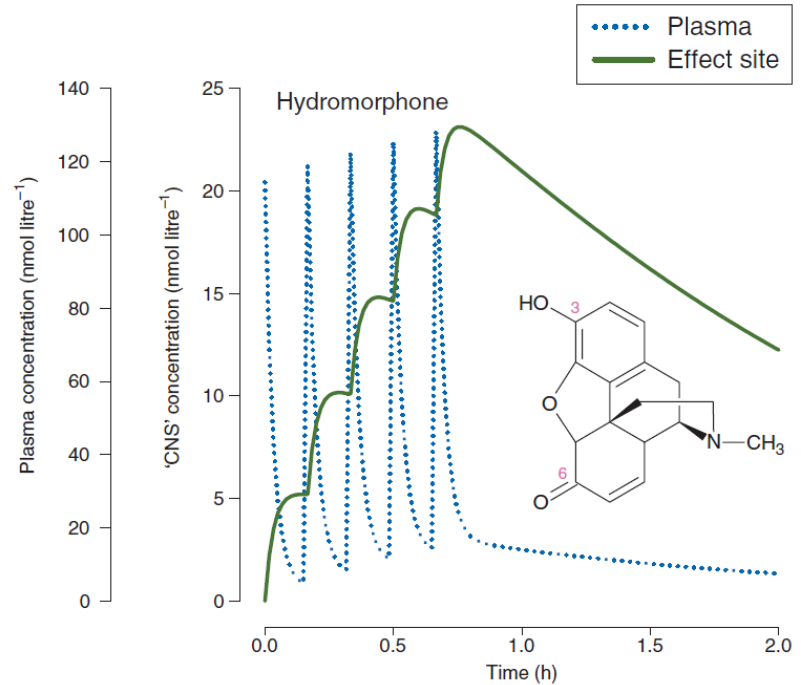
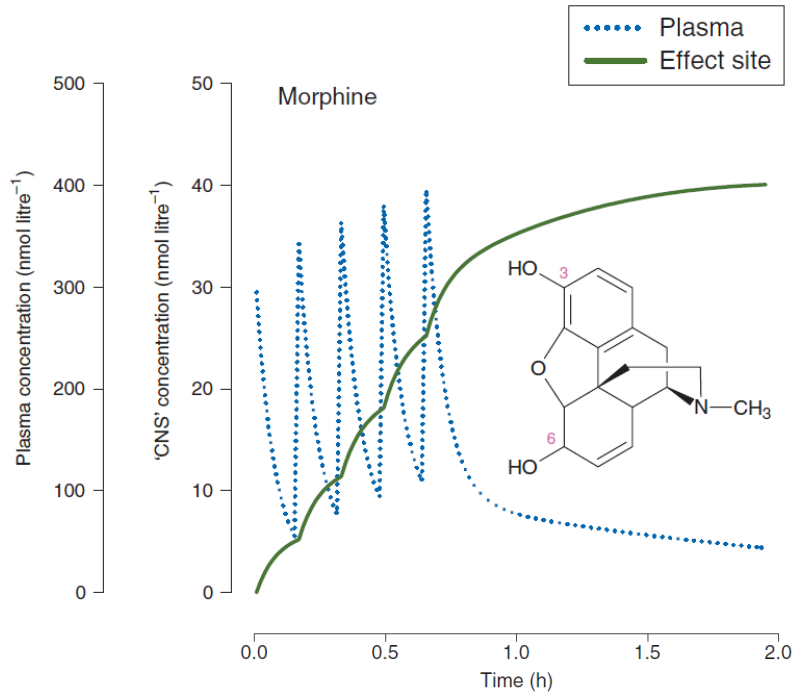
Das Verdünnen von Vendal<sup>®</sup>-Ampullen mit 0.9% NaCl oder 5%iger Glucoselösung ist laut Fachinformation möglich.

~~Siehe Fachinformation (Beilage) Punkt 6.2: Inkompatibilitäten.~~

**Die Injektionslösung darf mit keinen anderen Produkten vermischt werden.**

# Effect Site Concentration

## Vergleich Morphin – Hydromorphon



Felden L. et al.: BJA 2011; 107 (3): 319-328

## Wirkdauer

Dosis

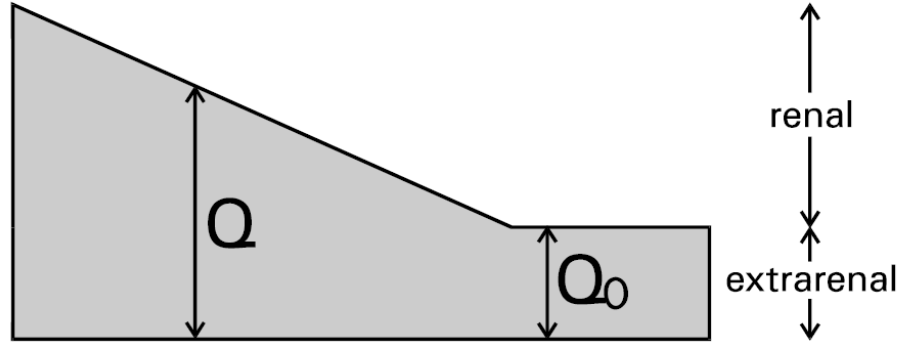
Halbwertszeit ( $T_{1/2}$ )

Verteilungsvolumen ( $V_T$ )

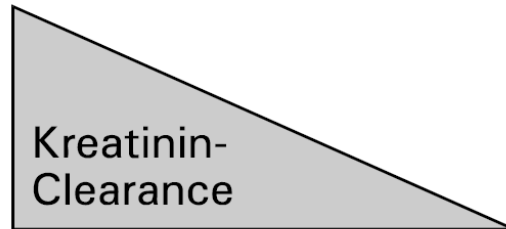
Kontextsensitive  
Halbwertszeit

# Extrarenale Dosisfraktion $Q_0$ bei Anurie renale Ausscheidung 0

$$Q_0 < 0,5$$



$$\text{Krea}_{\text{Cl}} < 50$$

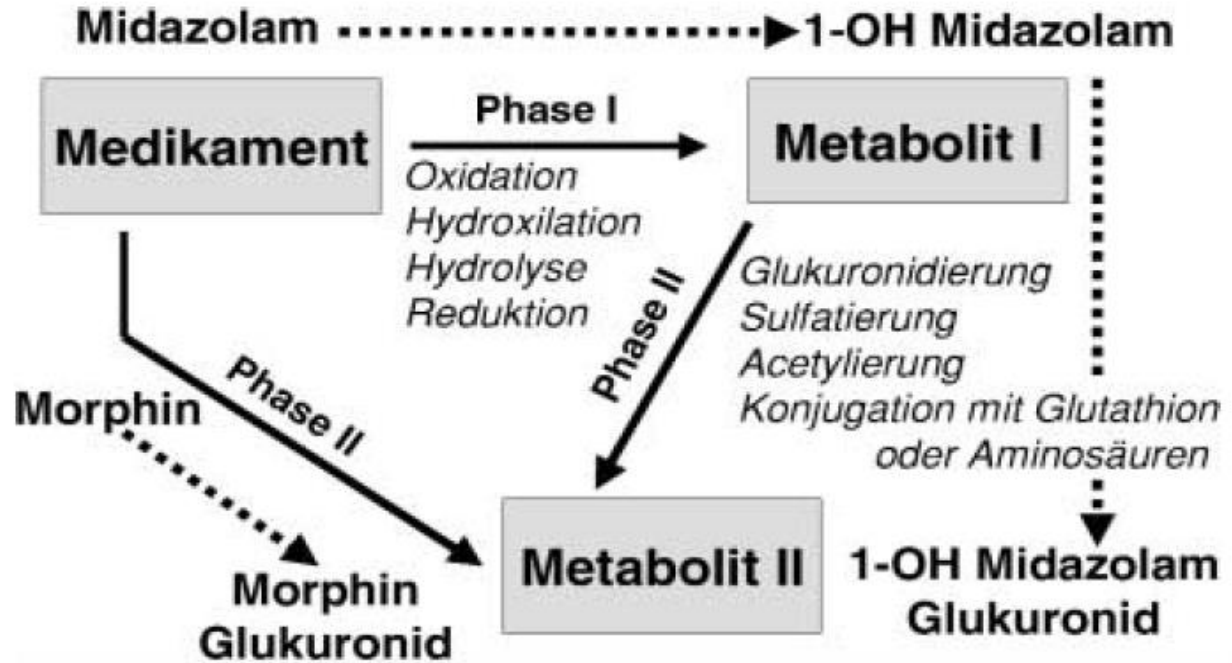


# Niereninsuffizienz

## EFIC

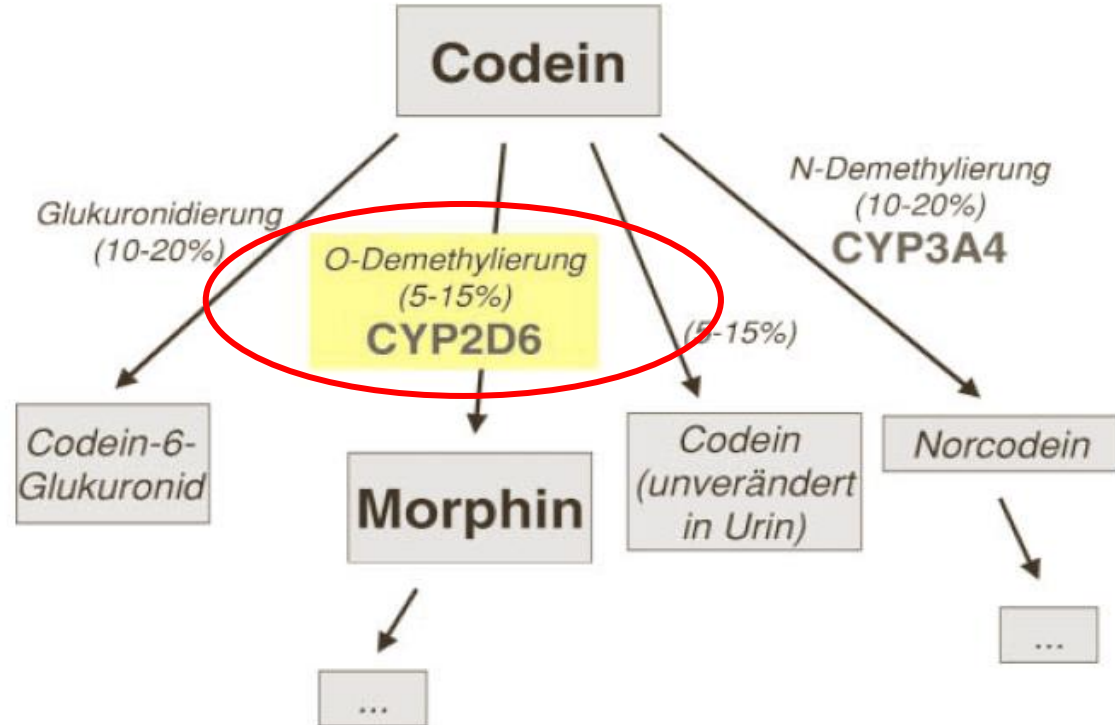
- empfohlen
  - Buprenorphin
  - Fentanyl
  - Oxymorphone
- „with caution“
  - Hydromorphone
  - Oxycodone
- should [...] not be prescribed“
  - Codeine
  - Dextropropoxyphene
  - Morphin
  - Pethidin
  - Tramadol

CYP abhängige und unabhängige hepatale Metabolisierung  
direkte Glucuronidierung ist weniger interaktionsgefährdet



# Elimination

unterschiedliche Abbauewege






aus Papp-Jambour et al. Anästhesist 2002; 51: 2-15



## Lebererkrankungen

### EFIC

- veränderte Pharmakokinetik
  - vermehrtes portocavales Shunting – First pass Effekt 
  - reduzierte Produktion von Transportproteinen – Plasmaspiegel 
  - verminderte metabolische Aktivität der Leber – hepatale Clearance 
- „only relevant in severe lever disease“
- Verlängerte Dosierungsintervalle, reduzierte Einzeldosen
  - Hydromorphon, Morphin, Oxycodon, Tapentadol, Tramadol
- Vermeiden
  - Codein, Pethidin

IONSYS®

Fentanyl Iontophoretic Transdermal System

Copyright ?

Actiq®

Instanyl®

Effentora®

Vellofent®

Copyright ?


## P2Y12 Inhibitoren

Hemmung der GP-IIb/IIIa-Aktivierung (ADP)

Copyright ?

SYSTEMATIC REVIEW

## The Comparative Risk of Delirium with Different Opioids: A Systematic Review

Lieke M. Swart<sup>1</sup>  · Vera van der Zanden<sup>1</sup> · Petra E. Spies<sup>2</sup> · Sophia E. de Rooij<sup>1,3</sup> ·  
Barbara C. van Munster<sup>2,3</sup>

There are no convincing data  
that the risk of delirium in elderly patients  
depends on the type of opioid.

	Biotransformation	Aktive Metaboliten	Q <sub>0</sub>	T <sub>1/2</sub> (h)
Tramadol (1970er, USA 1990er)	CYP 2D6 (stark) CYP 2B6, 3A4 (schwach)	O-Desmethyltramadol	~ 0,5	6
Nalbuphin	Glucuronidierung CYP 2C9, CYP 2C19	Nalbuphin-6-Glucuronid	hoch*	3
Piritramid (1960er)	CYP	k.A.	1,0*	7
Morphin (Anfang 20. Jh.)	Glucuronidierung	Morphin-6-Glucuronid (30%) M-3-G (60%, inaktiv)	0,9	2,4
Oxycodon (1916)	Glucuronidierung CYP 3A4/A5, CYP 2D6	Oxymorphon (19%, CYP 2D6) Noroxycodon (45%, inaktiv)	0,89	2,3
Methadon (1930er)	CYP3A4, CYP 2B6	Keine	k.A.	30 (7-130)
Hydromorphon (1921)	Glucuronidierung	Keine (Hydromorphon-3-Glucuronid neurotoxisch)	0,23	2,6
Burprenorphin (1977)	CYP 3A4, CYP 2C8 Glucuronidierung	Norbuprenorphin	1	3**
Fentanyl (1960)	CYP 3A4	Keine	0,9	3

\*keine Daten zum Einsatz bei Niereininsuffizienz

\*\* CAVE: langsame Dissoziation vom μ-Rezeptor

	Biotransformation	Aktive Metaboliten	$Q_0$	$T_{1/2}$ (h)
Tramadol (1970er, USA 1990er)	CYP 2D6 (stark) CYP 2B6, 3A4 (schwach)	O-Desmethyltramadol	~ 0,5	6
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Piritramid (1960er)	CYP	k.A.	1,0*	7
Morphin (Anfang 20. Jh.)	Glucuronidierung	<b>M-6-G:</b> - $T_{1/2}$ 2,8 h - $Q_0 \leq 0,3$ - $T_{1/2}$ bei Niereninsuffizienz bis 30 h		
Oxycodon (1916)	Glucuronidierung CYP 3A4/A5, CYP 2D6			
Methadon (1930er)	CYP3A4, CYP 2B6			
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\*\* CAVE: langsame Dissoziation vom  $\mu$ -Rezeptor

~~Buprenorphin~~

~~Methadon~~

~~Pethidin~~

Fentanyl

Morphin

Piritramid

Hydromorphon

Nalbuphin

~~Tapentadol~~

Oxycodon

Tramadol



		Wirkstärke (iv.)	Typ. Ampulle	
Tramadol		0,1	100 mg	(1 ml)
Nalbuphin		0,5	20 mg	(2 ml)
Piritramid		0,7	15 mg	(2 ml)
Morphin		1	10 mg	(1 ml)
Oxycodon		1	10 mg	(1 ml)
Diacetylmorphin		3	5 mg	(1 ml)
Hydromorphon		5	2 mg	(1 ml)
Burprenorphin		30	0.3 mg	(1 ml)
Fentanyl		100	0.1 mg	(2 ml)

## REDUCTION OF POSTOPERATIVE PAIN BY ENCOURAGEMENT AND INSTRUCTION OF PATIENTS\*

### A Study of Doctor-Patient Rapport

LAWRENCE D. EGBERT, M.D.,† GEORGE E. BATTIT, M.D.,‡ CLAUDE E. WELCH, M.D.,§  
AND MARSHALL K. BARTLETT, M.D.¶

BOSTON

**M**ANY reports have discussed the treatment of patients suffering after operation. Narcotics are not without danger; they also vary considerably in effectiveness. Hypnosis will reduce pain but is difficult to achieve and requires special training for the operator. Despite considerable effort the problems of treating postoperative pain remain.

Janis<sup>1</sup> has shown that patients who were told about their operations before the procedure remembered the operation and its sequelae more favorably than those who were not well informed. We have determined the effects of instruction, suggestion and encouragement upon the severity of postoperative pain.

#### METHOD

the following day, they were told what would be done about the pain. They were advised that pain is caused by spasm of the muscles under the incision and that they could relieve most of the pain themselves by relaxing these muscles. They could achieve relaxation by slowly taking a deep breath and consciously allowing the abdominal wall to relax. Also, they were shown the use of a trapeze that was hanging over the middle of the bed (control patients also had the trapeze but were not instructed by the anesthetist). Special-care patients were taught how to turn onto one side by using their arms and legs while relaxing their abdominal muscles. Finally, they were told that at first they would find it difficult to relax completely. If they

Egbert L. et al.: NEJM 1964; 270 (16): 825-827

## REDUCTION OF POSTOPERATIVE PAIN AND INSULIN REQUIREMENT

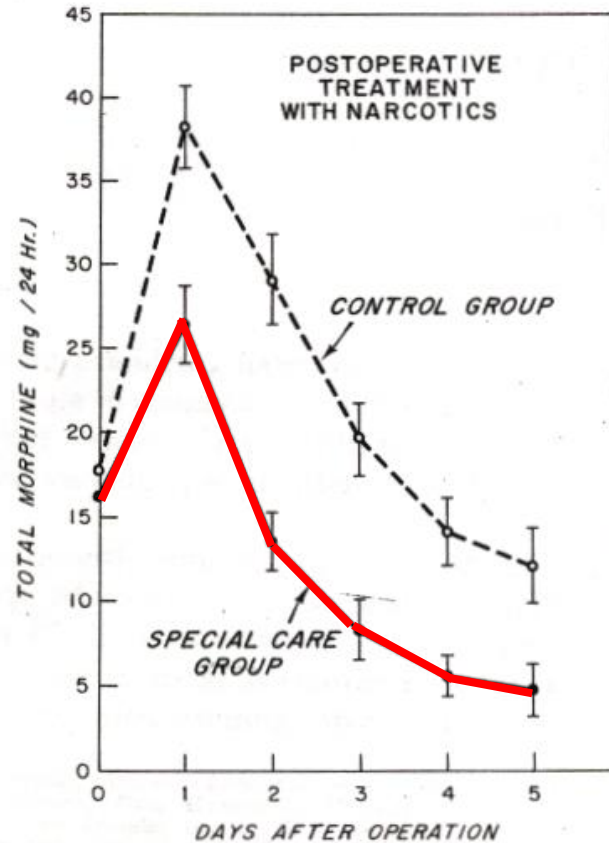
A Study

LAWRENCE D. EGBERT, M.D.,†  
AND M. J. JANIS, M.D.

MANY reports have discussed the treatment of patients suffering after operation. These treatments are not without danger; they also vary considerably in effectiveness. Hypnosis will reduce pain, but is difficult to achieve and requires special training of the operator. Despite considerable effort the methods of treating postoperative pain remain.

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### METHOD



Egbert L. et al.: NEJM 1964; 270 (16): 825-827



Prof. Dr. Andrea Michalek-Sauberer



Priv. Doz. Dr. Sylvia Reichl