#### BfS Workshop May 9-10, 2007

#### Effects of HF Signals on the Melatonin Synthesis in Isolated Pineal Organs of Djungarian Hamsters

(Untersuchungen zu Wirkungsmechanismen an Zellen unter Exposition mit hochfrequenten elektromagnetischen Feldern der Mobilfunktechnologie. B. Pinealdrüse BfS StSch 4318)

> Alexander Lerchl Jacobs University Bremen

# 1800 MHz electromagnetic field effects on melatonin release from isolated pineal glands

Abstract: Isolated pineal glands of Djungarian hamsters (*Phodopus sungorus*) were continuously perifused by Krebs-Ringer buffer, stimulated with the beta-adrenergic receptor agonist isoproterenol to induce melatonin synthesis, and exposed for 7 hr to a 1800 MHz continuous wave (CW) or pulsed GSM (Global System for Mobile Communications)-modulated electromagnetic signal at specific absorption rate (SAR) rates of 8, 80, 800, and 2700 mW/kg. Experiments were performed in a blind fashion. Perifusate samples were collected every hour, and melatonin concentrations were measured by a specific radioimmunoassay. Both types of signal significantly enhanced melatonin release at 800 mW/kg SAR, while at 2700 mW/kg SAR, melatonin levels were elevated in the CW, but suppressed in the GSM-exposed pineal glands. As a temperature rise of approximately 1.2°C was measured at 2700 mW/kg SAR, effects at this level are thermal. With regard to radiofrequency electromagnetic fields, the data do not support the 'melatonin hypothesis,' according to which nonthermal exposure suppresses melatonin synthesis.

#### Irina Sukhotina<sup>1,2</sup>, Joachim R. Streckert<sup>3</sup>, Andreas K. Bitz<sup>3</sup>, Volkert W. Hansen<sup>3</sup> and Alexander Lerchl<sup>1</sup>

<sup>1</sup>School of Engineering and Science, International University Bremen, Bremen, Germany; <sup>2</sup>Valdman Institute of Pharmacology, Pavlov Medical University, St Petersburg, Russia; <sup>3</sup>Chair of Electromagnetic Theory, University of Wuppertal, Wuppertal, Germany

Key words: electromagnetic fields, hamsters, melatonin, mobile phone, pineal gland

Address reprint requests to Alexander Lerchl, International University Bremen, School of

# Introduction

In the discussion about the possible mechanisms that are involved in the biologic effects of ELF-MF or RF-EMF, the <u>melatonin hypothesis</u> plays a central role, based on experimental data from the late 1970s showing reduced melatonin values after exposure to electric or magnetic fields.

Original Model:

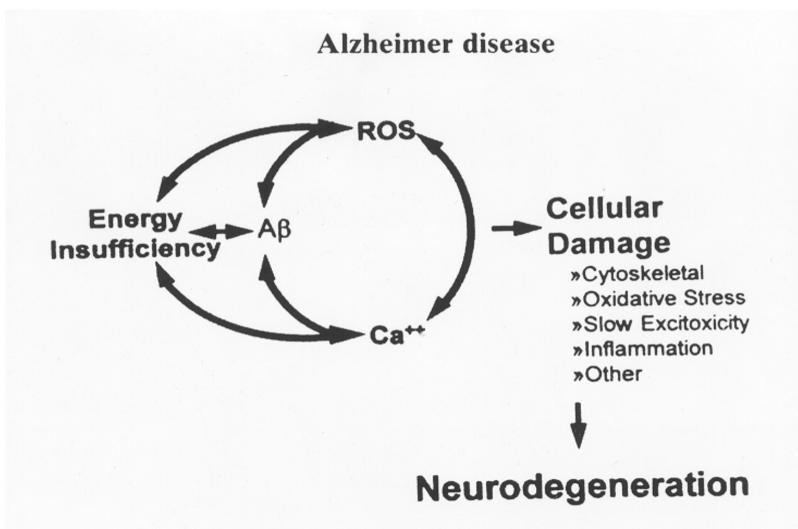
ELF MF/EF  $\rightarrow$  low melatonin  $\rightarrow$  high estrogens  $\rightarrow$  increased breast tumor growth

# Introduction

In the mean time (since the early 1990s) it became clear that melatonin has in fact some properties which may be relevant for many diseases.

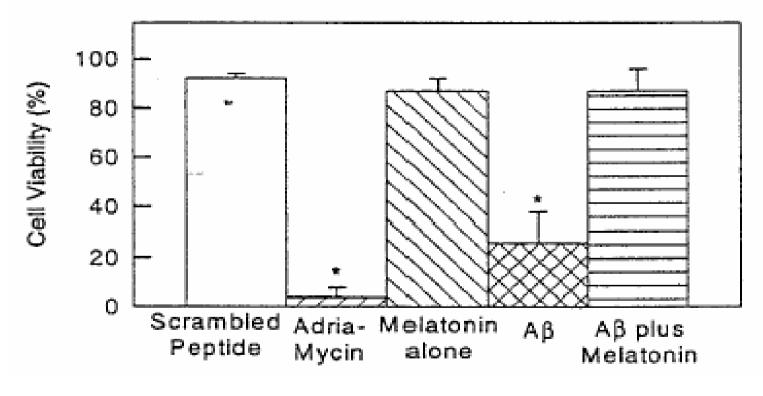
- -Cancer
- -Stroke (reperfusion)
- -Coronary infarction
- -and other diseases which are caused by / related to ROS

#### Melatonin is a Radical Scavenger for Reactive Oxygen Species (ROS)



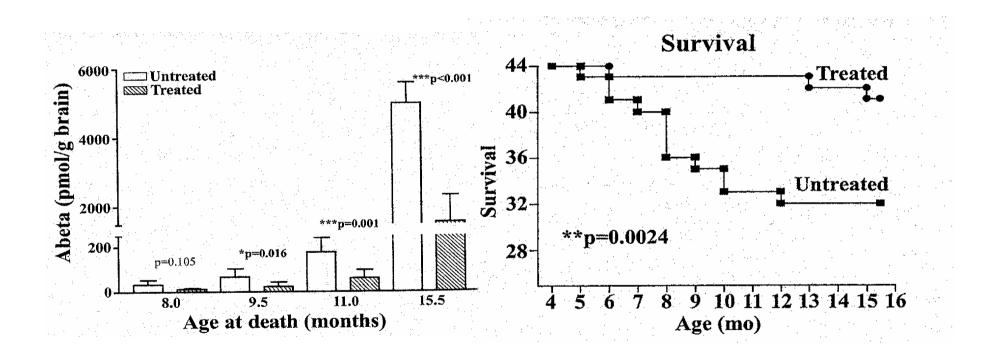
#### Melatonin and Alzheimer's Disease

Effects of the Amyloid-B Peptide on Cell Survival in Neuroblastoma Cells



Pappolla et al., 1997

#### Prevention of Alzheimer's Disease by Melatonin in Transgenic Mice



Matsubara et al., 2003

#### **RF-EMF** effects on Melatonin

Hamster / Rat	EMF, 900 MHz, 0,04-0,36 W/Kg	15 min-6 hrs	Melatonin +/-	Vollrath et al. 1997
Rat	EMF, 900 and 1800 MHz, 100 / 20 μW/cm²	2 hrs per day, 14 days	6-OHMS +/-	Bakos et al., 2003
Human	Mobile phone use	2 weeks, questionaire	6-OHMS -	Burch et al., 2002
Human	EMF, 900 MHz, 1 W/m²	4 hrs	Melatonin +/-	Radon et al. 2001
Human	900 and 1800 MHz	2 hrs / day, 5 days / week, 4 weeks	Melatonin +/-	De Seze et al., 1999
Human	EMF, 900 MHz (GSM)	night	Melatonin +/-	Mann et al. 1998

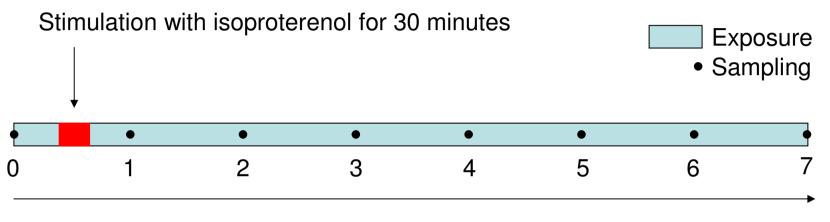
# Aims of the Study

Relatively little is known about the effect of RF-EMF on pineal function. In the present study, we investigated the <u>direct</u> effects of high frequency 1800 MHz electromagnetic fields of 8, 80, 800, and 2700 mW/kg SAR levels on melatonin release by isolated pineal organs from Djungarian hamsters. Both CW and pulsed signals were used. The pulse signals were modulated according to the GSM.

# Methods

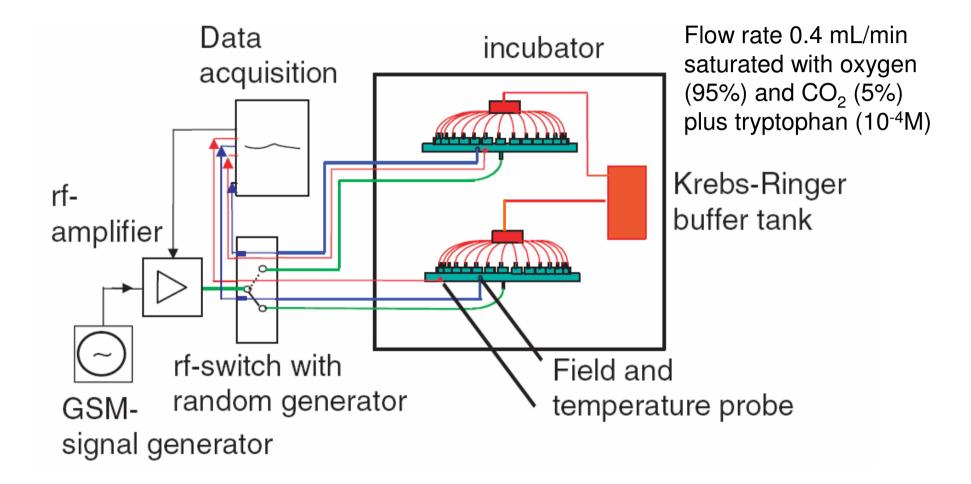
- n=20 pineal organs per experiment (based on statistical pre-analyses: 25% difference detectable at α=0.05 and B=0.80)
- Sham-exposed and exposed (shielding better than -70dB), blind design
- Incubation for 7 hrs, sampling every hour
- Stimulation with isoproterenol for 30 min.
- Measurement of melatonin production by specific radioimmunoassay
- Calculation of pg melatonin mL<sup>-1</sup> hr<sup>-1</sup>

# Methods

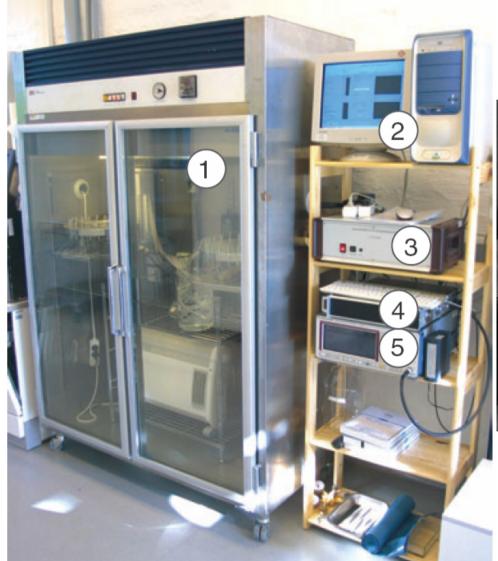


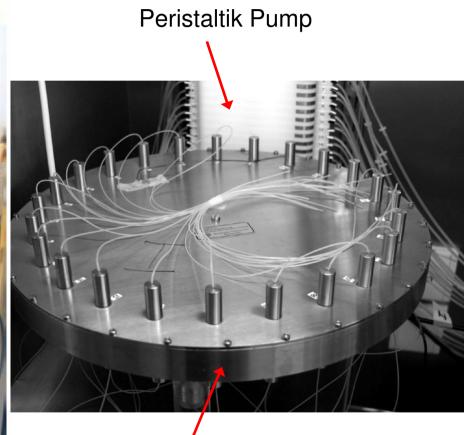
Time (Hours)

#### Methods: Overview



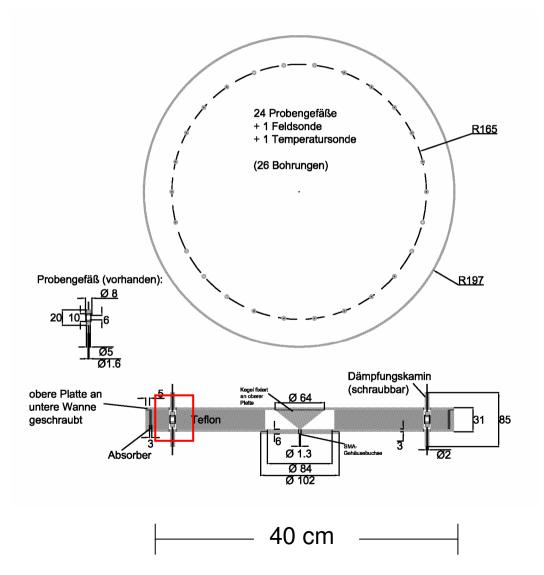
#### Exposure

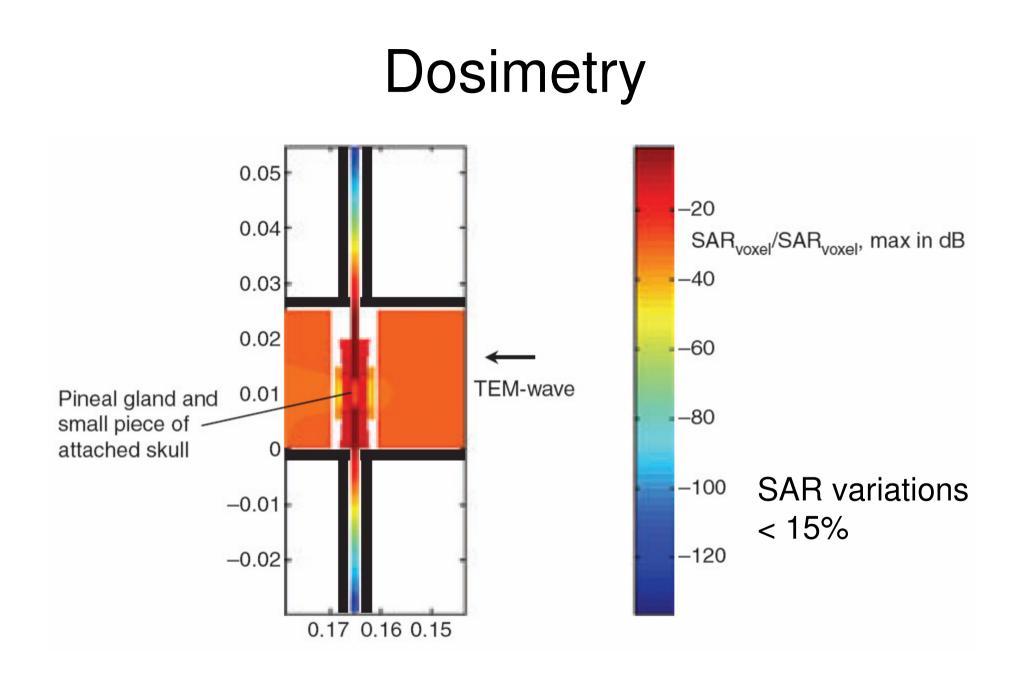




Radial Waveguide

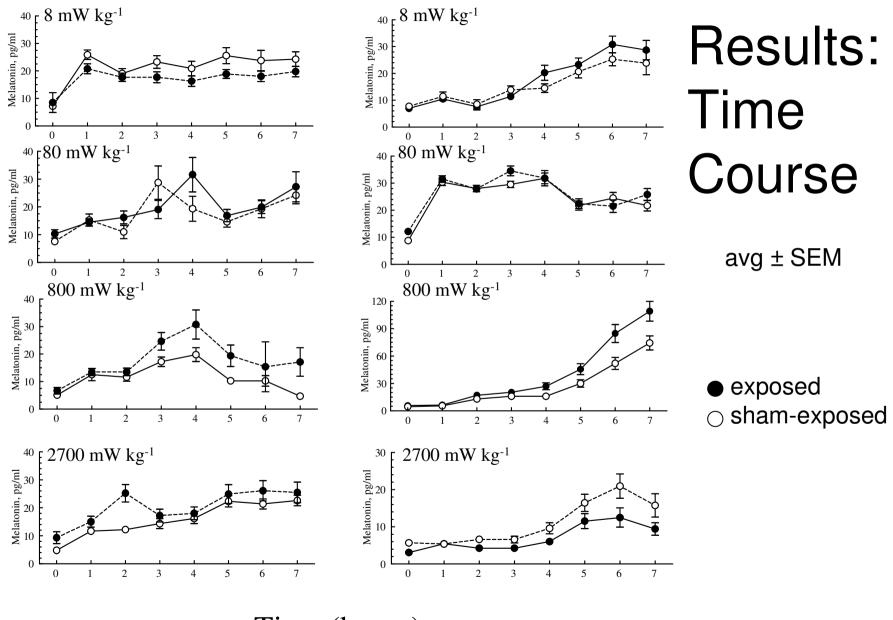
#### **Radial Waveguides**





CW

GSM



Time (hours)

#### **Results: Table**

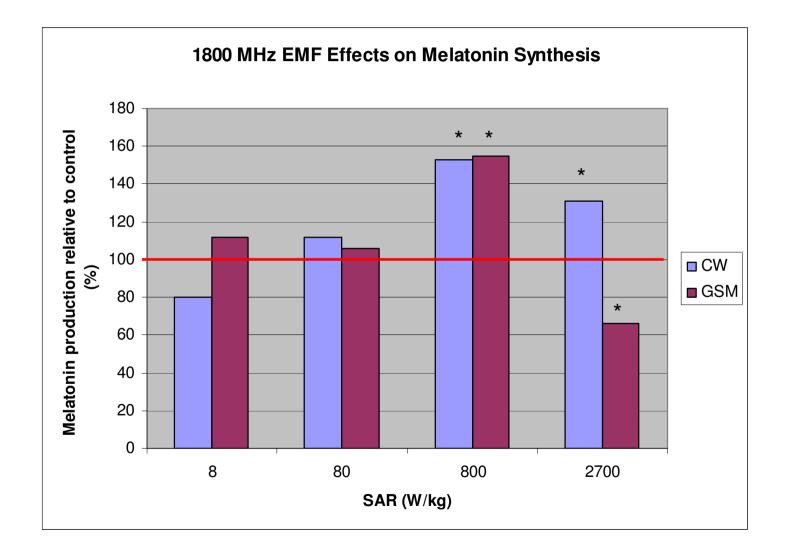
	Dose (mW/kg)				
Melatonin production	8	80	800	2700	
CW					
In control pineal glands	$21.75 \pm 0.29$	$17.24 \pm 0.31$	$11.47 \pm 0.28$	$15.65 \pm 0.28$	
In exposed pineal glands	$17.22 \pm 0.30$	$19.31 \pm 0.31$	$17.54 \pm 0.32$	$20.45 \pm 0.29$	
Relative to control (%)	79.8	112.0	152.9*	130.7*	
GSM					
In control pineal glands	$15.65 \pm 0.30$	$24.62 \pm 0.29$	$25.65 \pm 0.38$	$10.44 \pm 0.28$	
In exposed pineal glands	$17.49 \pm 0.31$	$26.04 \pm 0.30$	$39.75 \pm 0.23$	$6.9~\pm~0.24$	
Relative to control (%)	111.8	105.8	155.0*	66.1*	

CW = continuous (unmodulated) exposure, GSM = modulated exposure according to the GSM standard.

Data represent melatonin synthesis  $(pg/mL/hr) \pm S.E.M.$ 

\*P < 0.05 versus control. For further statistical results see text.

#### **Results:** Figure



# Discussion

- No effects were seen at 8 and 80 mW/kg SAR.
- Temperature rises at 2700 mW/kg were approximately 1.2°C, thus thermal effects were likely to have caused the effects.
- At 800 mW/kg SAR, a temperature rise of approximately 0.5 – 0.6 ℃ due to exposure may have contributed to the elevated melatonin levels.

# Conclusions

- At SAR levels relevant to human exposure of the pineal organ\*, no adverse effects on melatonin synthesis (i.e. decreases) were observed.
- These data therefore do not support the melatonin hypothesis for RF-EMF.

\*maximum calculated average SAR levels <0.1 W/kg in childrens' heads; Bit-Babik et al., 2005

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