



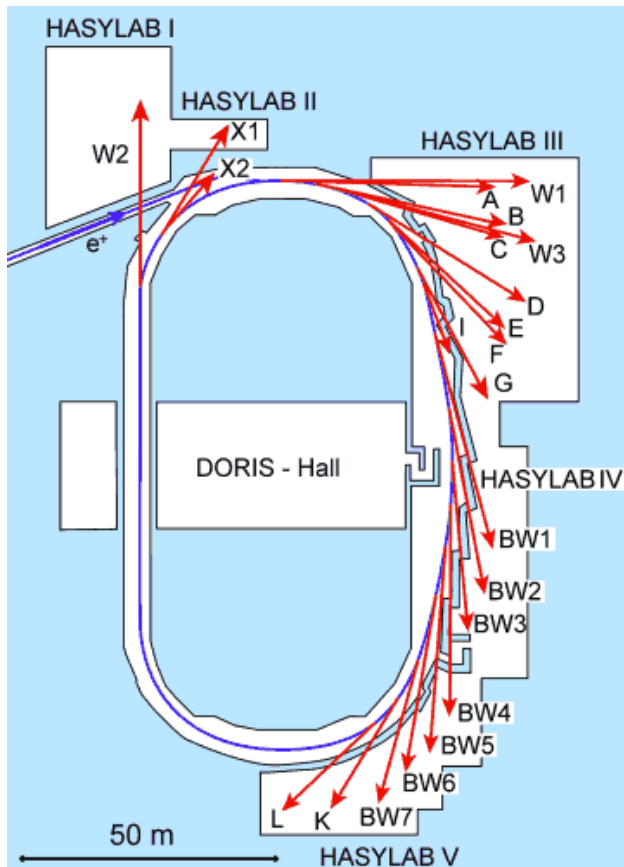
# Tomographic imaging using synchrotron light: Benefit for orthopedic research

Frank Witte





# Synchrotron-radiation based computed micro-tomography



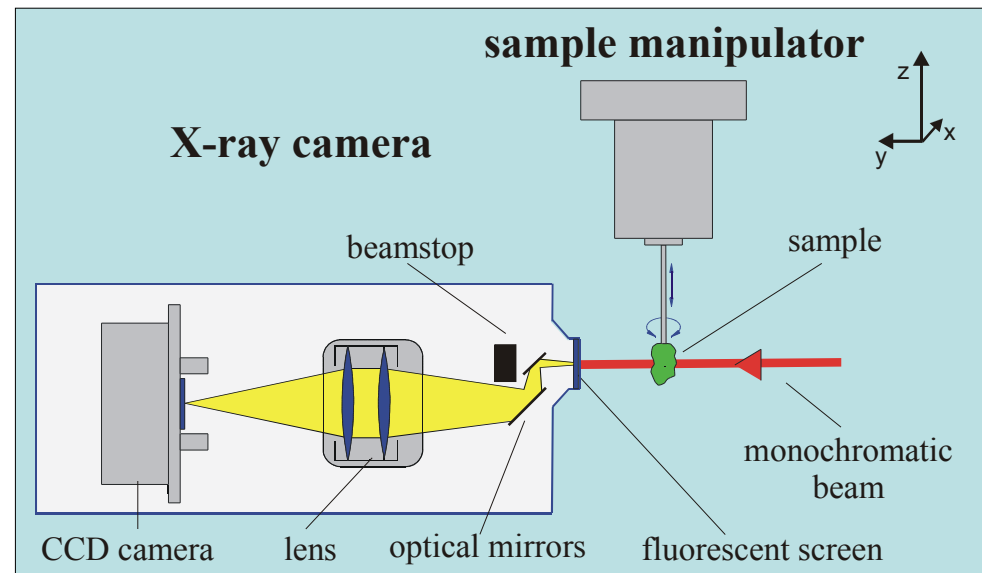
## Hamburger Synchrotronstrahlungslabor

### HASYLAB

- DORIS storage ring
- $E=4.450$  GeV Positrons
- 22 Beamlines



HARWI II  
Felix Beckmann





# The beginning of SR $\mu$ CT in biology

644

Nuclear Instruments and Methods in Physics Research A246 (1986) 644–648  
North-Holland, Amsterdam

## HIGH RESOLUTION TOMOGRAPHY WITH CHEMICAL SPECIFICITY

U. BONSE <sup>2)</sup>, Q. JOHNSON <sup>1)</sup>, M. NICHOLS <sup>3)</sup>, R. NUSSHARDT <sup>2)</sup>, S. KRASNICKI <sup>2)</sup>\*  
and J. KINNEY <sup>1)</sup>

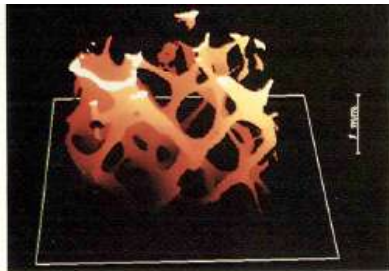
<sup>1)</sup> Lawrence Livermore National Laboratory, Livermore, CA 94550, USA

<sup>2)</sup> Institut für Physik, University of Dortmund, Dortmund, FRG

<sup>3)</sup> Sandia National Laboratory, Livermore, CA 94550, USA



Prof. Ulrich Bonse



Bone and Mineral 25 (1994) 25–38

**BONE AND MINERAL**

### 3D computed X-ray tomography of human cancellous bone at 8 $\mu$ m spatial and $10^{-4}$ energy resolution

Ulrich Bonse\*<sup>a</sup>, Frank Busch<sup>a</sup>, Olaf Günnewig<sup>a</sup>, Felix Beckmann<sup>a</sup>,  
Reinhard Pahl<sup>a</sup>, Günter Delling<sup>b</sup>, Michael Hahn<sup>b</sup>, Walter Graeff<sup>c</sup>

<sup>a</sup>Lehrstuhl für Experimentelle Physik I, Universität Dortmund, Otto-Hahn-Str. 4, D-44221 Dortmund, Germany

<sup>b</sup>Institut für Pathologie, Universitäts-Krankenhaus Eppendorf, Martinistr. 52, 20240 Hamburg, Germany

<sup>c</sup>Hamburger Synchrotronstrahlungslabor HASYLAB at Deutsches Elektronensynchrotron DESY, Notkestr. 85, 22607 Hamburg, Germany

(Received 19 August 1993; revision received 28 October 1993; accepted 10 November 1993)



Microcallus (www.roche.com)

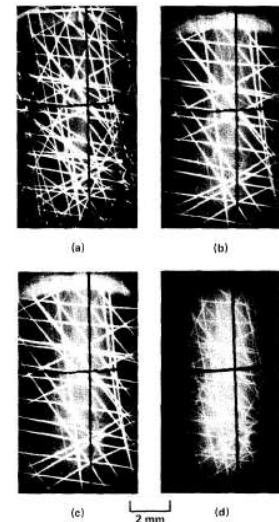


Fig. 7. (a) Web structured gaps in Cu+Ni Layers imaged at 23 eV above Cu K-edge, (b) 18 eV below Cu K-edge, (c) 26 eV above Ni K-edge, and (d) 22 eV below Ni K-edge.





# SR $\mu$ CT-based FEM



Journal of Biomechanics ■ (■■■■) ■■■-■■■

JOURNAL  
OF  
BIOMECHANICS

[www.elsevier.com/locate/jbiomech](http://www.elsevier.com/locate/jbiomech)  
[www.JBiomech.com](http://www.JBiomech.com)

## Intrinsic mechanical properties of trabecular calcaneus determined by finite-element models using 3D synchrotron microtomography

H. Follet<sup>a,\*</sup>, F. Peyrin<sup>b,c</sup>, E. Vidal-Salle<sup>a</sup>, A. Bonnassie<sup>b</sup>, C. Rumelhart<sup>a</sup>, P.J. Meunier<sup>d</sup>

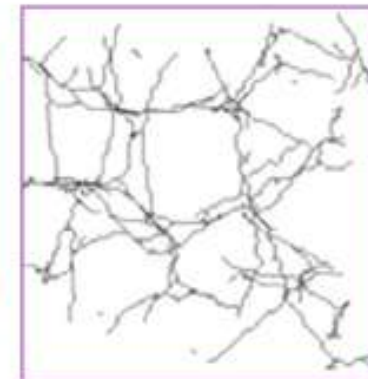
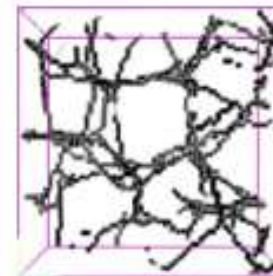
<sup>a</sup>Laboratoire de Mécanique des Contacts et des Solides (LaMCoS) UMR CNRS 5514, INSA, Bât Coulomb, Lyon, France

<sup>b</sup>CREATIS, UMR CNRS 5515, Bât. Blaise Pascal, INSA, Lyon, France

<sup>c</sup>ESRF, BP 220, 38043 Grenoble Cedex, France

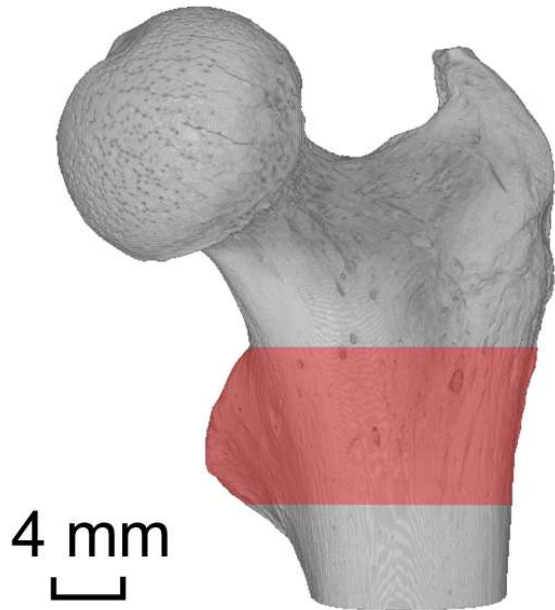
<sup>d</sup>Laboratoire d'Histodynamique Osseuse, INSERM U403, Lyon, France

Accepted 24 October 2006

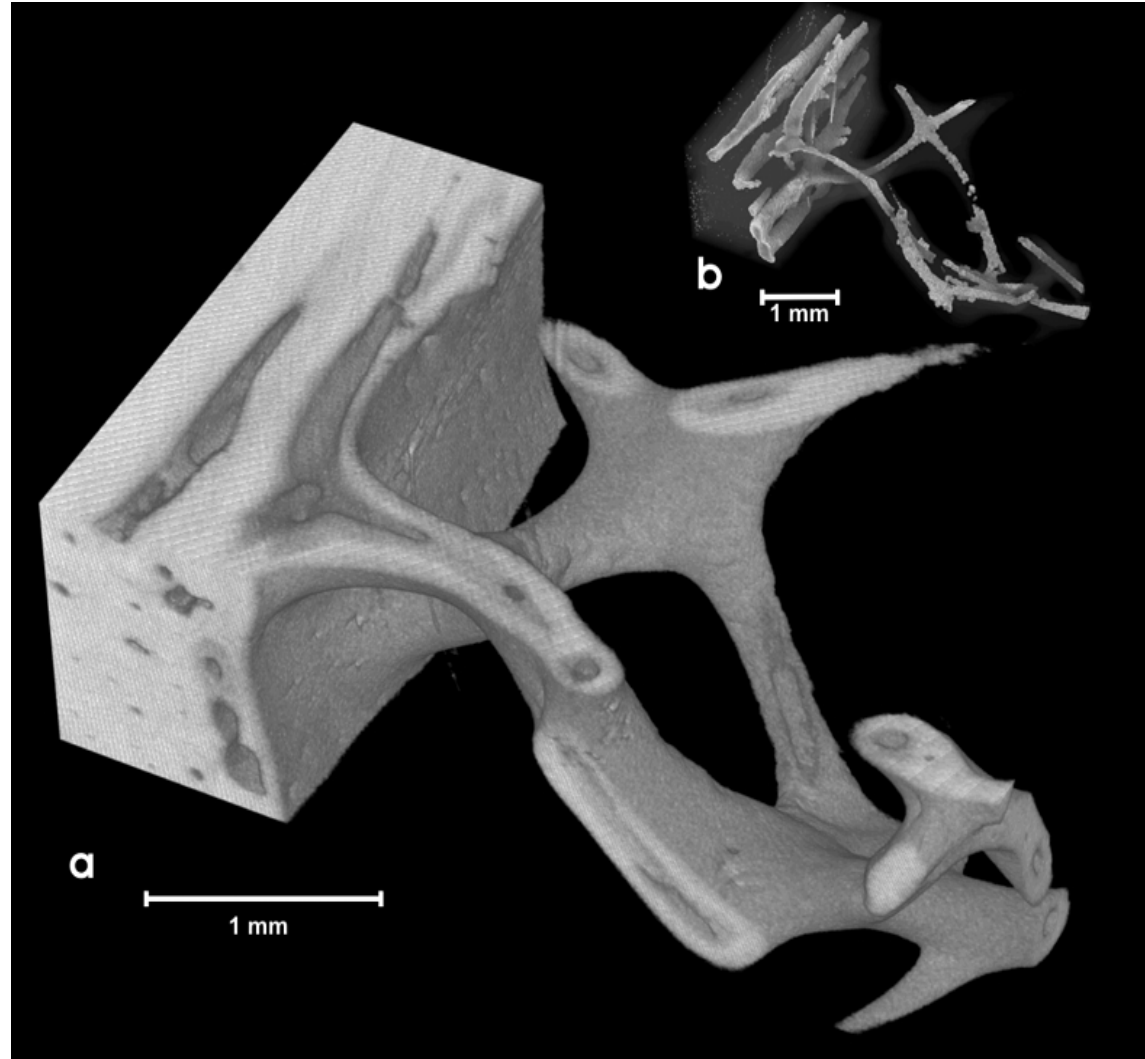




# SR $\mu$ CT for structural biology



Scherf H., Fischer J., Beckmann F.,  
Witte F. SPIE 5535, 2004







# SR $\mu$ CT for soft tissue imaging

Phys. Med. Biol. 43 (1998) 2911–2923. Printed in the UK

PII: S0031-9155(98)93345-0

*Orthod Craniofacial Res* 9, 2006/199–205

ORIGINAL ARTICLE

*M Dalstra  
PM Cattaneo  
F Beckmann*

Synchrotron radiation-based  
microtomography of alveolar  
support tissues

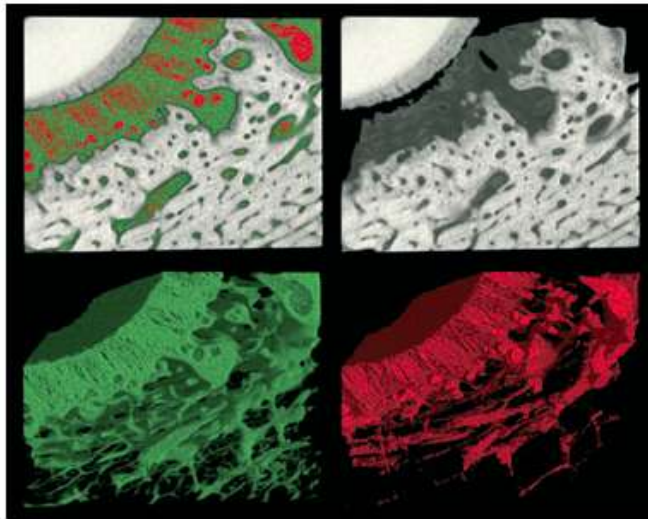


Fig. 8. 3D reconstruction of the mineralized and soft tissues in a section of the porcine sample. Note the use of false-coloring to enhance the low-density structures; green for the bone marrow and the fibers of the PDL; red for the blood vessels and interstitial tissue.

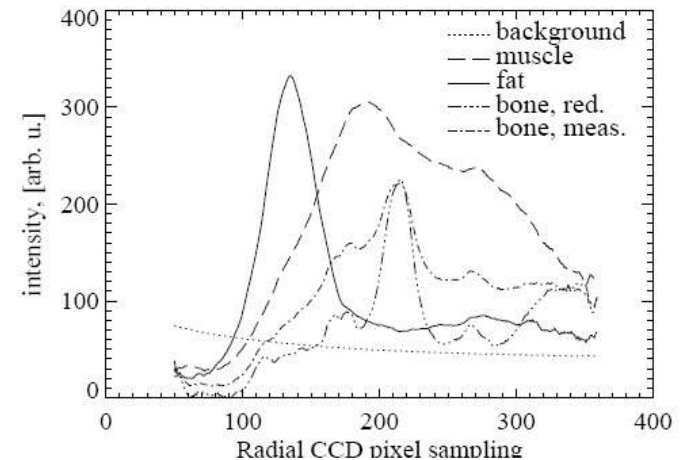
## Feasibility study of x-ray diffraction computed tomography for medical imaging\*

U Kleuker<sup>†‡</sup>, P Suortti<sup>†</sup>, W Weyrich<sup>§</sup> and P Spanne<sup>†</sup>

<sup>†</sup> Medical Imaging Group, European Synchrotron Radiation Facility, PO Box 220, F-38043 Grenoble, France

<sup>§</sup> Faculty of Chemistry, University of Konstanz, Fach M721, D-78457 Konstanz, Germany

Received 16 April 1998, in final form 27 July 1998





# SR $\mu$ CT in tissue engineering

Microsc. Microanal. 12, 97–105, 2006  
DOI: 10.1017/S1431927606060168

Microscopy AND  
Microanalysis

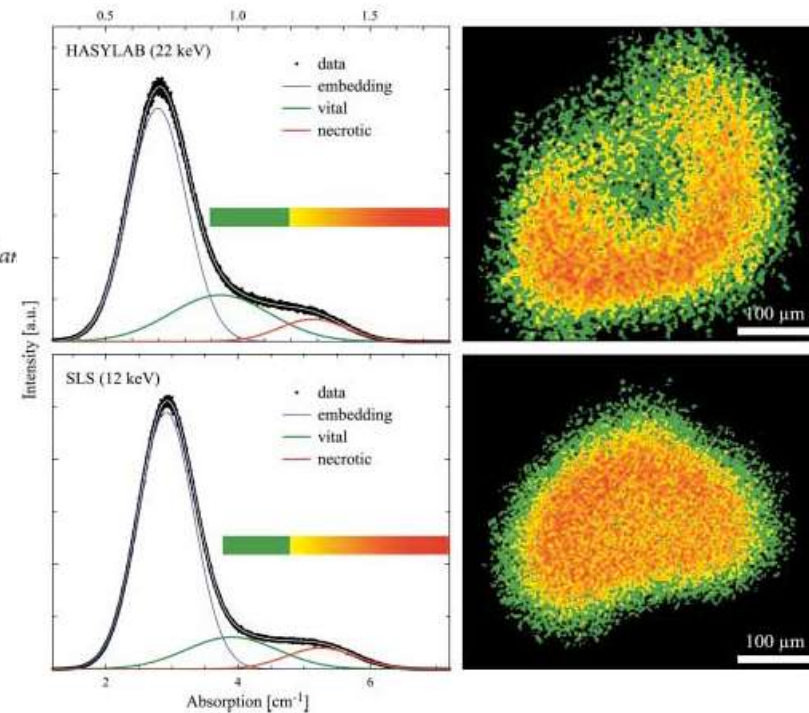
## Three-Dimensional Characterization of Cell Clusters Using Synchrotron-Radiation-Based Micro-Computed Tomography

Bert Müller,<sup>1,\*</sup> Marco Riedel,<sup>2</sup> Philipp J. Thurner<sup>1,3</sup>

<sup>1</sup>Computer Vision Laboratory ETH Zürich, Gloriastrasse 35, CH-8092 Zürich, Switzerland

<sup>2</sup>ProBioGen, Sternwartstrasse 7, D-13086 Berlin, Germany

<sup>3</sup>Swiss Federal Institute for Materials Testing and Research, Überlandstrasse 129, CH-8600 Dübendorf, Switzerland





# Biomaterial research using SR $\mu$ CT

- Field of application
  - Structural bone analysis
  - Bone-Implant-Interface
  - Determination of degradation rates
- Mainly used modes
  - SR $\mu$ CT: attenuation mode
  - SR $\mu$ CT: element-specific mode







# Limitation of hard-tissue histology

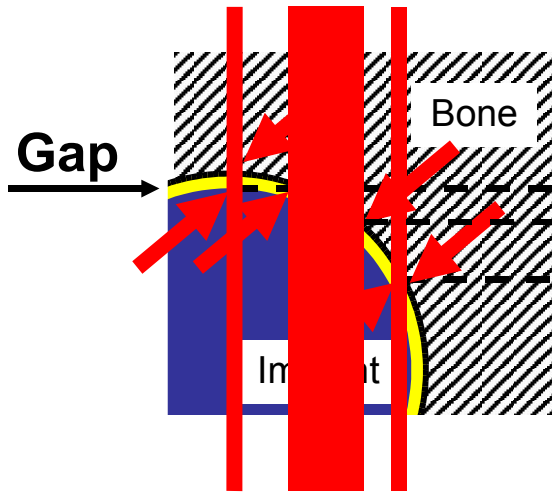
1. Limitation of *biomaterial-tissue interface* analysis
  - „Wrong angle“ phenomena
  - limitation of cutting-grinding technique



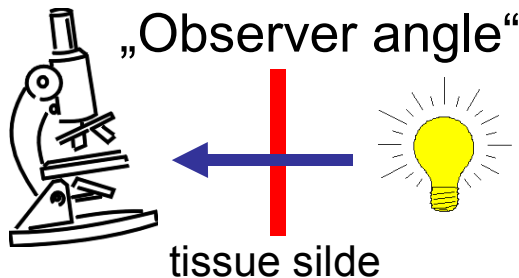
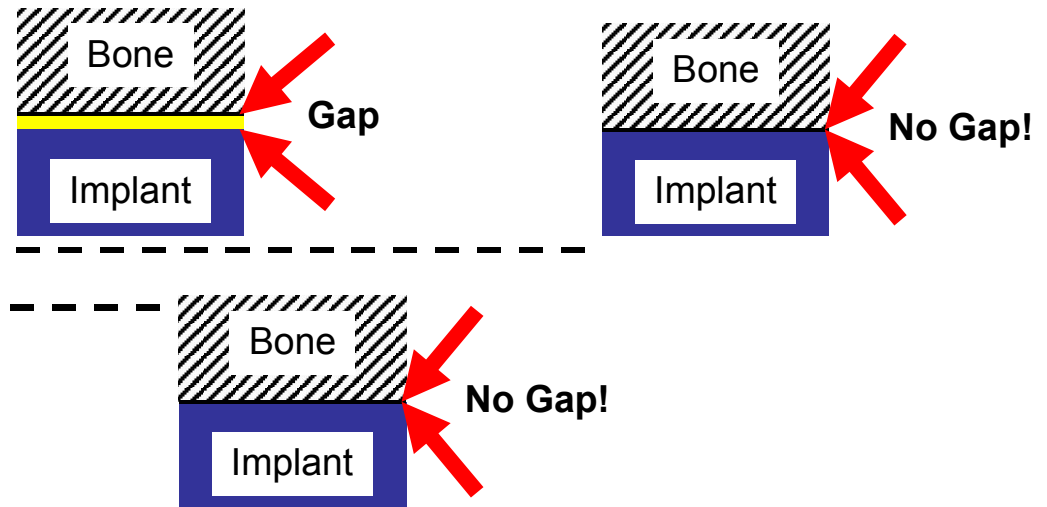


# „Wrong angle“ – Phenomena

Plane of Histological sections



Appearance of Bone-Implant-Interphase



„wrong-angle“-phenomena is dependent on:

1. thickness of the histological section
2. the plane of the section (An and Friedman, 1999)





# Advantage of SR $\mu$ CT

European Cells and Materials Vol. 7. 2004 (pages 42-51)

ISSN 1473-2262

## COMPARISON OF MICROFOCUS- AND SYNCHROTRON X-RAY TOMOGRAPHY FOR THE ANALYSIS OF OSTEOINTEGRATION AROUND TI6AL4V-IMPLANTS

R. Bernhardt<sup>1</sup>, D. Scharnweber<sup>1</sup>, B. Müller<sup>2\*</sup>, P. Thurner<sup>3</sup>, H. Schliephake<sup>4</sup>, P. Wyss<sup>3</sup>, F. Beckmann<sup>5</sup>, J. Goebbels<sup>6</sup>, and H. Worch<sup>1</sup>

<sup>1</sup>Max-Bergmann-Center of Biomaterials, TU Dresden, D-01062 Dresden, Germany,

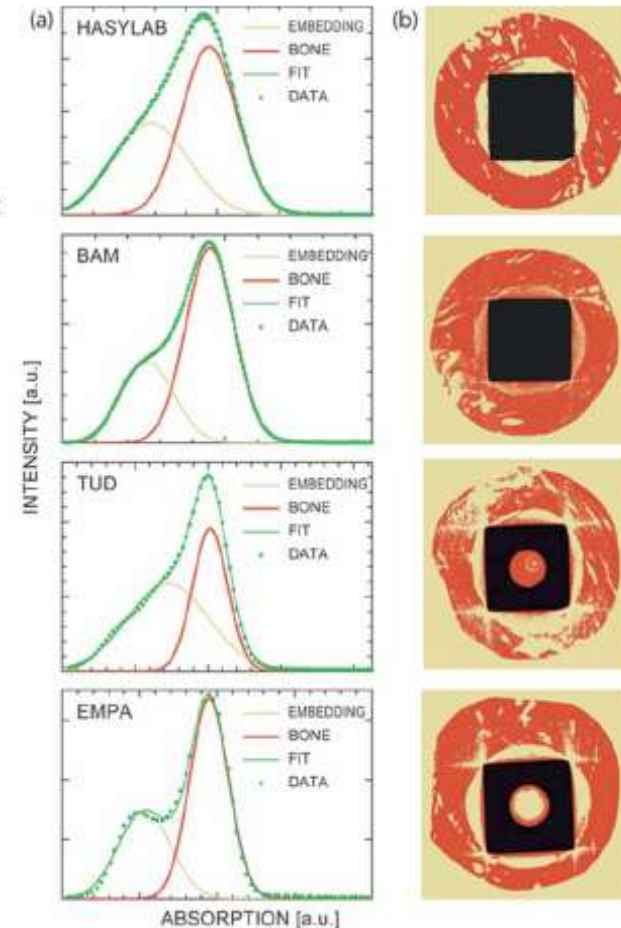
<sup>2</sup>Computer Vision Laboratory, Swiss Federal Institute of Technology, CH-8092 Zürich, Switzerland

<sup>3</sup>Swiss Federal Laboratories for Materials Testing and Research, CH-8600 Dübendorf, Switzerland

<sup>4</sup>Department of Cranio-Maxillofacial Surgery, Georgia-Augusta-University, D-37075 Göttingen, Germany

<sup>5</sup>GKSS-Research Center, D-21502 Geesthacht, Germany

<sup>6</sup>Federal Institute for Materials Research and Testing, D-12200 Berlin, Germany





# SR $\mu$ CT Interface Analysis

## Influence of extracellular matrix coatings on implant stability and osseointegration: An animal study

Bernd Stadlinger 1 \*, Eckart Pilling 1, Matthias Huhle 1, Ronald Mai 1, Susanne Bierbaum 2, Ricardo Bernhardt 2, Dieter Scharnweber 2, Eberhard Kuhlisch 3, Ute Hempel 4, Uwe Eckelt 1

1Faculty of Medicine, Department of Maxillofacial Surgery, University of Technology Dresden, Fetscherstr. 74, D-01307 Dresden, Germany

2Max-Bergmann-Center of Biomaterials, University of Technology, Budapester Str. 27, D-01062 Dresden, Germany

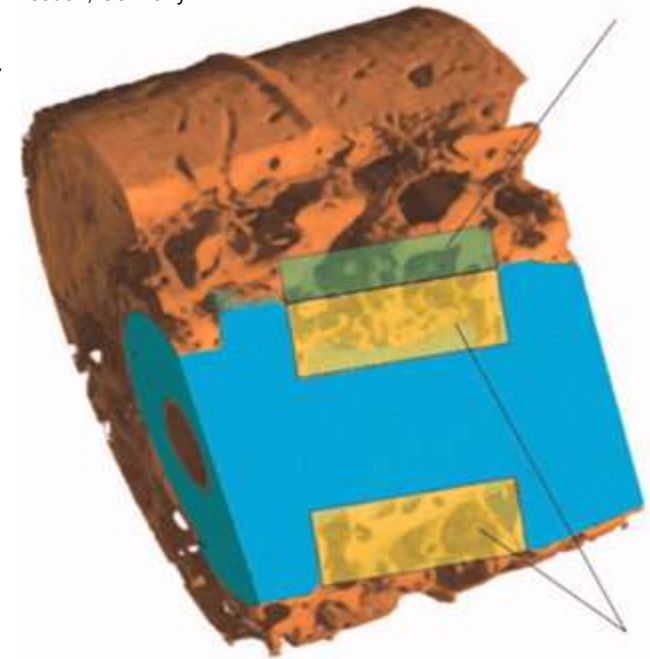
3Faculty of Medicine, Institute for Medical Informatics and Biometry, University of Technology Dresden, Fetscherstr. 74, D-01307 Dresden, Germany

4Faculty of Medicine, Institute of Physiological Chemistry, University of Technology Dresden, Fetscherstr. 74, D-01307 Dresden, Germany

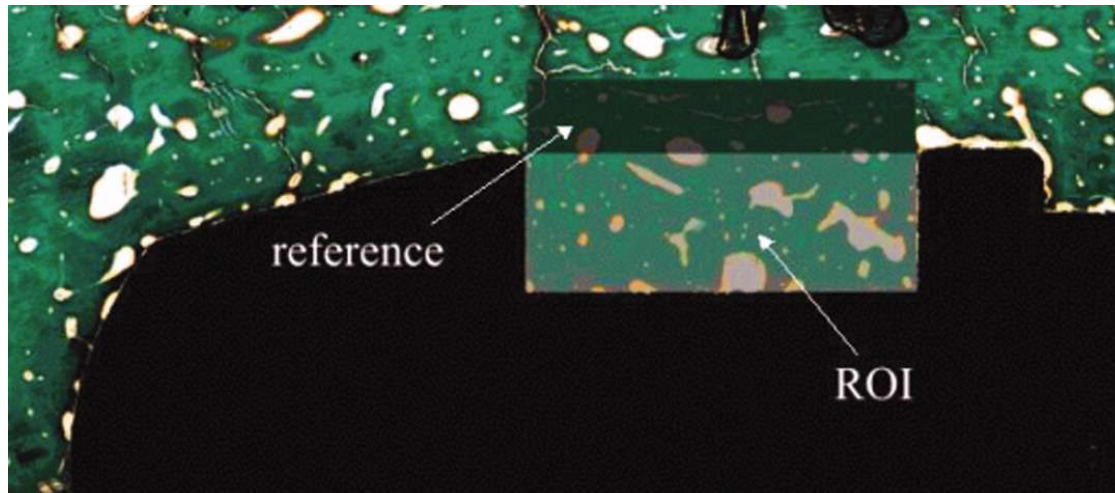
email: Bernd Stadlinger ([stadlinger@gmx.de](mailto:stadlinger@gmx.de))

**Journal of Biomedical Materials Research Part B: Applied Biomaterials**; Epub 22.Feb.2007

reference



ROI



reference

ROI







# SR $\mu$ CT Interface Analysis



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Biomaterials 26 (2005) 3009–3019

Biomaterials

[www.elsevier.com/locate/biomaterials](http://www.elsevier.com/locate/biomaterials)

## Osteoconductive modifications of Ti-implants in a goat defect model: characterization of bone growth with SR $\mu$ CT and histology

Ricardo Bernhardt<sup>a,\*</sup>, Juliette van den Dolder<sup>b</sup>, Sussane Bierbaum<sup>a</sup>, Rene Beutner<sup>a</sup>, Dieter Scharnweber<sup>a</sup>, John Jansen<sup>b</sup>, Felix Beckmann<sup>c</sup>, Hartmut Worch<sup>a</sup>

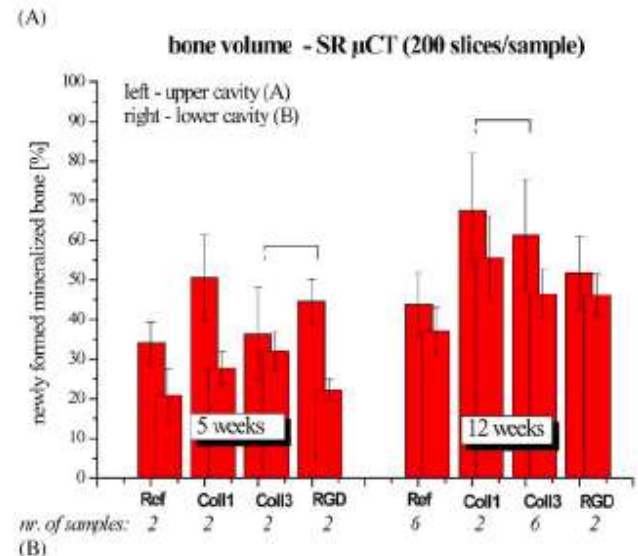
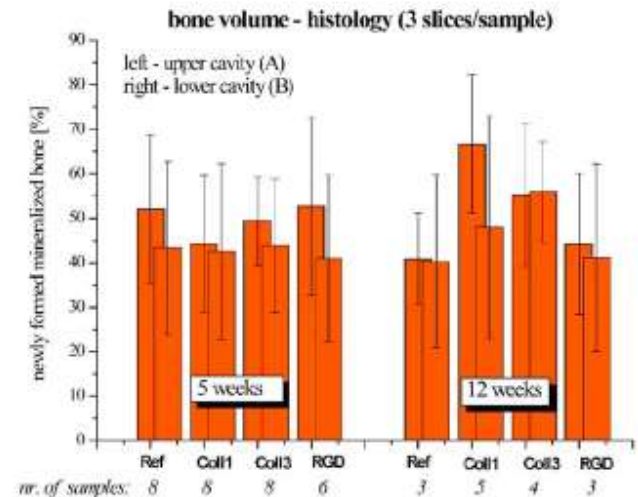
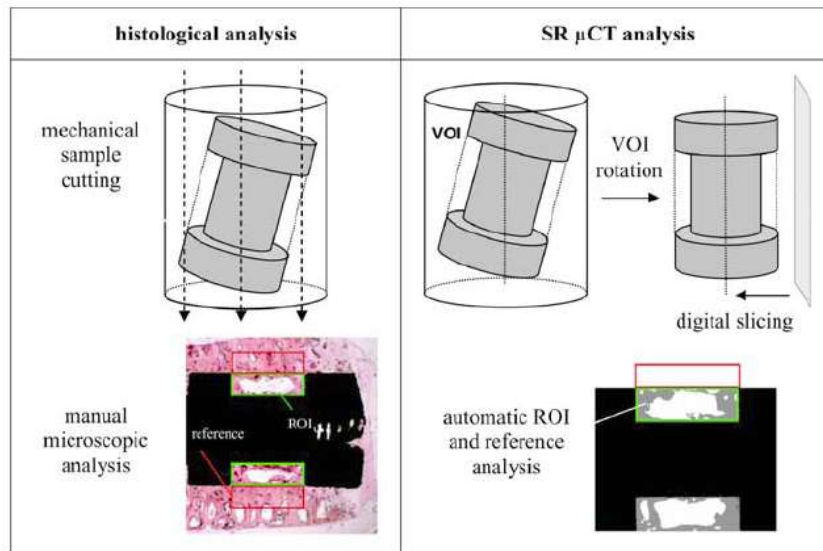
<sup>a</sup>Max-Bergmann-Center of Biomaterials, Dresden, University of Technology, Budapest Str. 27, D-01062 Dresden, Germany

<sup>b</sup>Department of Biomaterials, University Medical Center, P.O. Box 9101, 6500 HB Nijmegen, The Netherlands

<sup>c</sup>GKSS-Research Center, D-21502 Geesthacht, Germany

Received 12 May 2004; accepted 26 August 2004

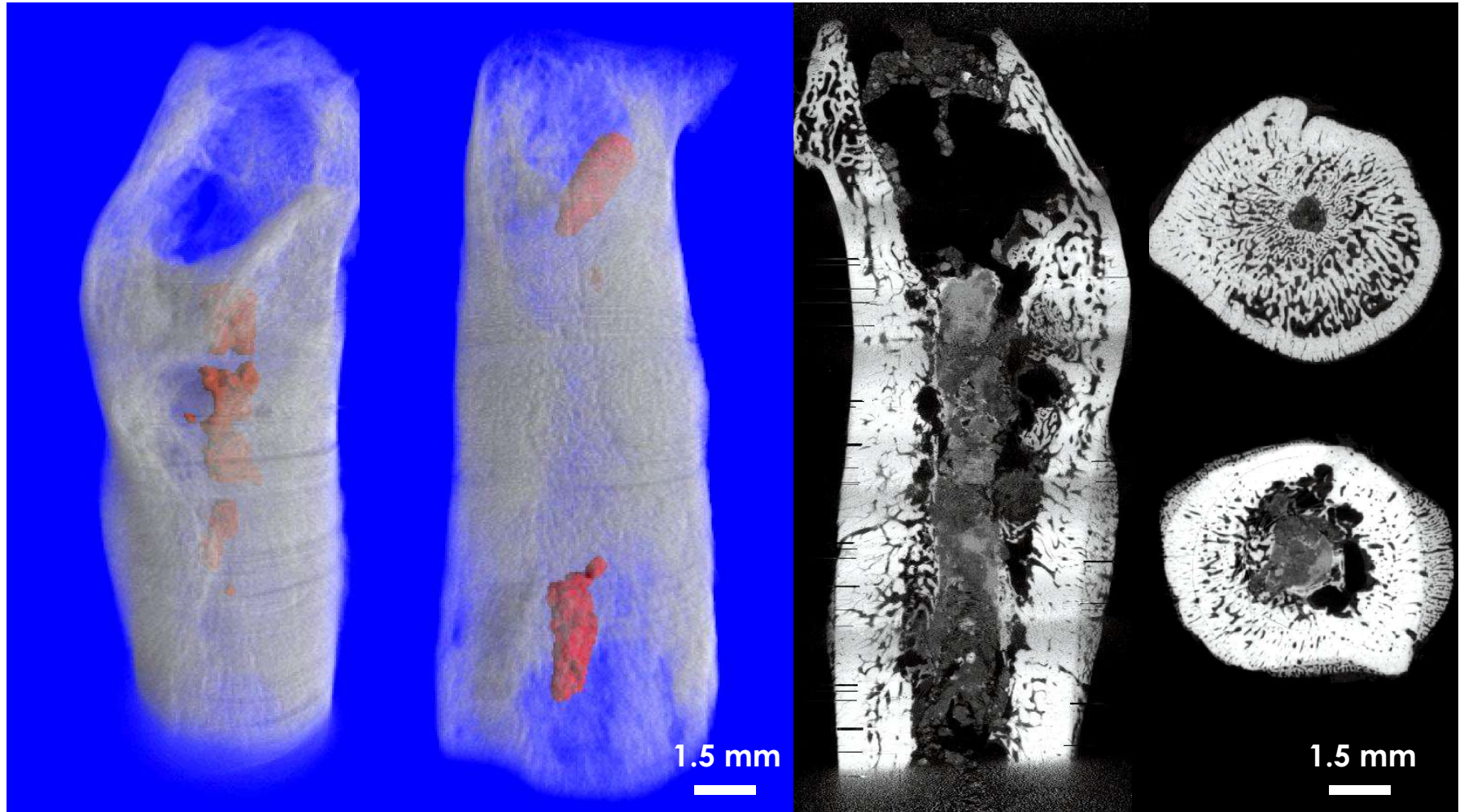
Available online 5 October 2004







# Biodegradable magnesium alloys

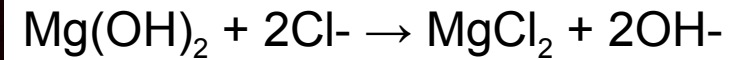
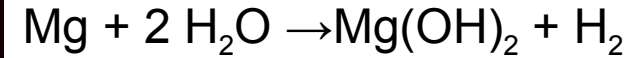
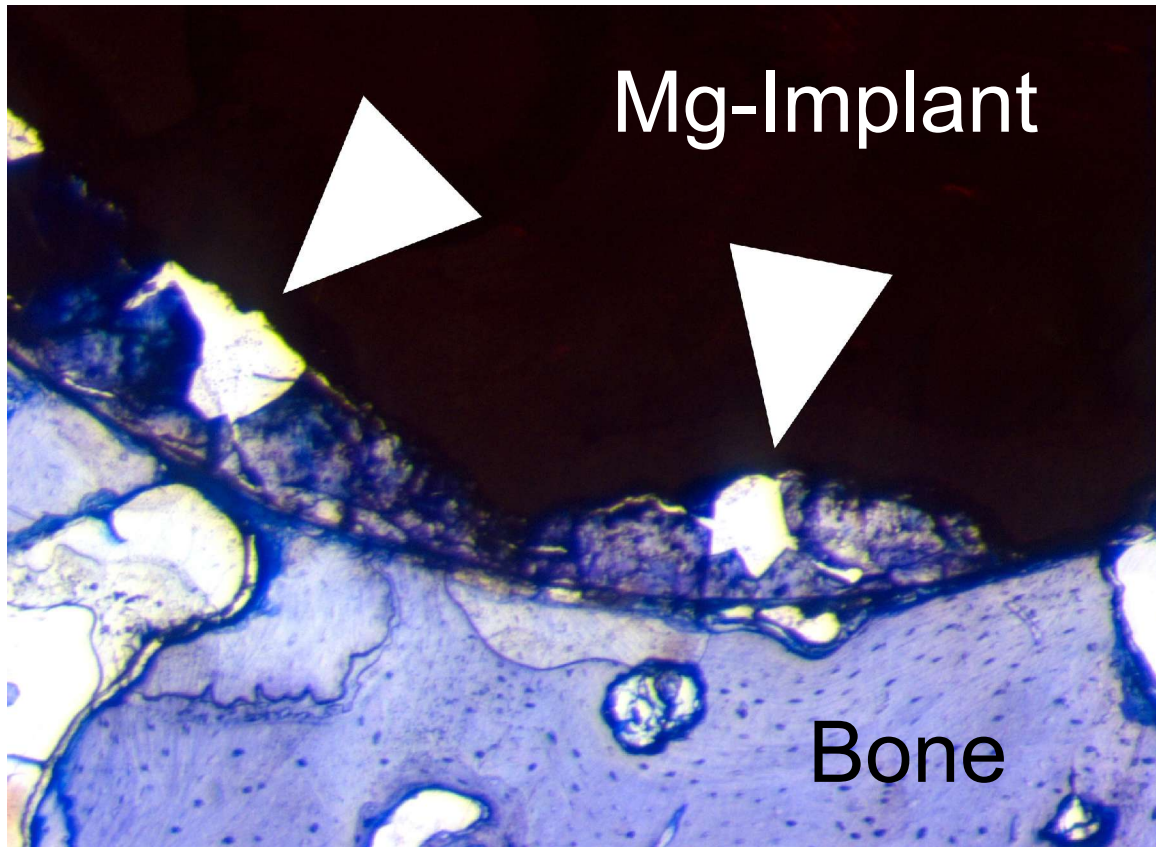


Witte et al., HASYLAB Ann. Reports, 2004





# Limitation of cutting-grinding technique



For analysis of in-vivo corrosion of magnesium, microtomography is mandatory.

*“Drop-outs”* from cutting-grinding technique





# Current biomaterials

material	density [g/cm <sup>3</sup> ]	Young's modulus [GPa]	tensile strength [Mpa]	breaking elongation [%]
surgical steel (X2CrNiMo18164)	8,0	193	585	55
surgical titanium (TiAl6V4)	4,43	100 – 110	930 – 1140	8 – 15
cortical bone	1,7 – 2,0	3 – 30	80 – 150	3 – 4
DL-PLA (DL-poly lactide)	1,24	1,9	29	5,0
magnesium AZ91	1,81	45	240	3
magnesium MgCa (0,8 wt% Ca)	1,75	n/a	290 – 300	2 – 6

Staiger et al., Biomaterials 27 2006





# Pathophysiology

0

2

## Body content of Mg = 21-35 g

**Bone** ↔ **Serum** ↔ **Muscle**  
**60%**                      **const. 1%**                      **39%**

**excess Mg**

**Kidney/Intestine**

1 H 1,00794	2 He 4,00260
3 Li 6,941	4 Be 9,012182
11 Na 22,989770	12 Mg 24,3050
19 K 39,0983	20 Ca 40,078
37 Rb 85,4678	38 Sr 87,62
55 Cs 132,90545	56 Ba 137,327
87 Fr (223)	88 Ra (226)







# Implantation of Magnesium-Rods



ELSEVIER

In vivo corrosion of four magnesium alloys and the associated bone response

F. Witte<sup>a,\*</sup>, V. Kaese<sup>b</sup>, H. Haferkamp<sup>b</sup>, E. Switzer<sup>c</sup>, A. Meyer-Lindenberg<sup>c</sup>,  
C.J. Wirth<sup>a</sup>, H. Windhagen<sup>a</sup>

<sup>a</sup>Department of Orthopaedic Surgery, Hannover Medical School, Anna-von-Borries-Str.1-7, 30625 Hannover, Germany

<sup>b</sup>Center of Biomedical Engineering, University of Hannover, Lise-Meitner-Str. 1, 30823 Garbsen, Germany

<sup>c</sup>Small Animal Clinic, School of Veterinary Medicine Hannover, Bischofsholer Damm 15, 30173 Hannover, Germany

Received 13 June 2004; accepted 22 September 2004

Available online 2 November 2004

**Biomaterials**

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Biomaterials 26 (2005) 3557–3563

Magnesium alloy:

AZ31

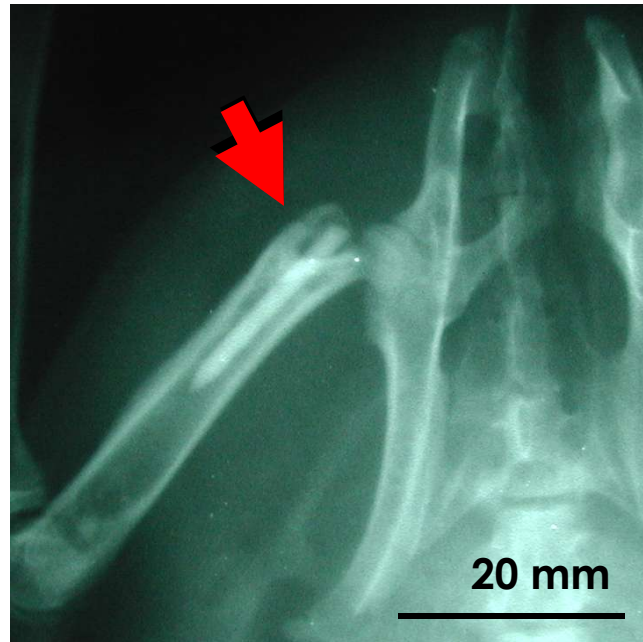
AZ91D

WE43

LAE442

Control:

SR-PLA



Follow-up post-OP:

6 weeks

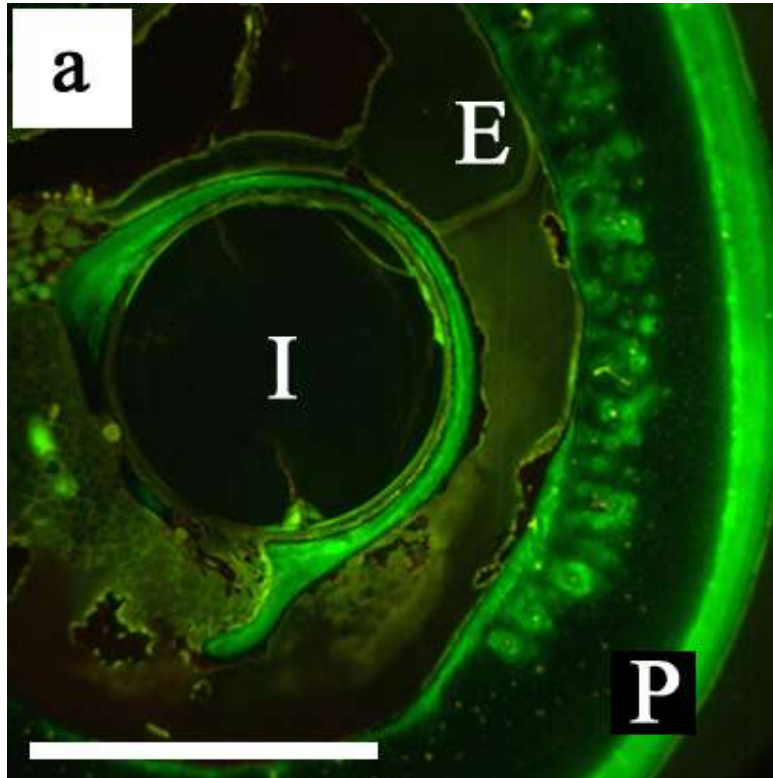
18 weeks



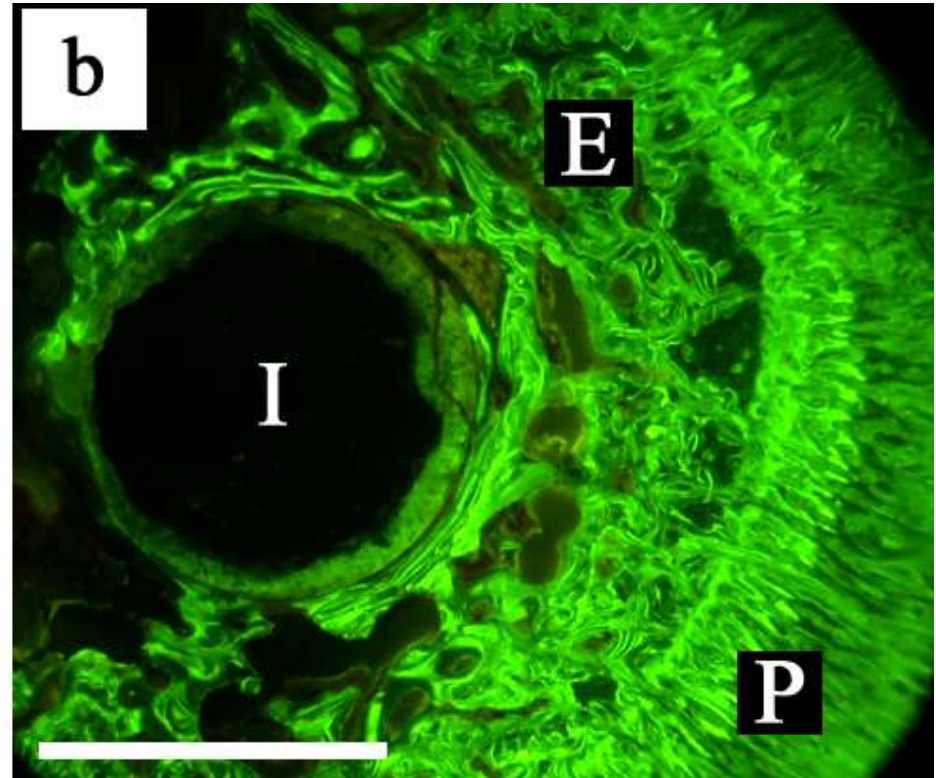




# New Bone Formation



SR-PLA96



Magnesium

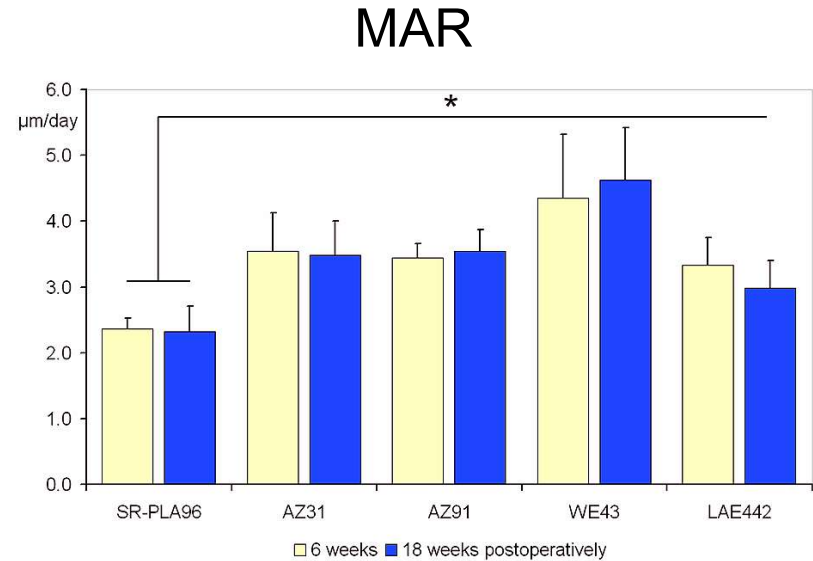
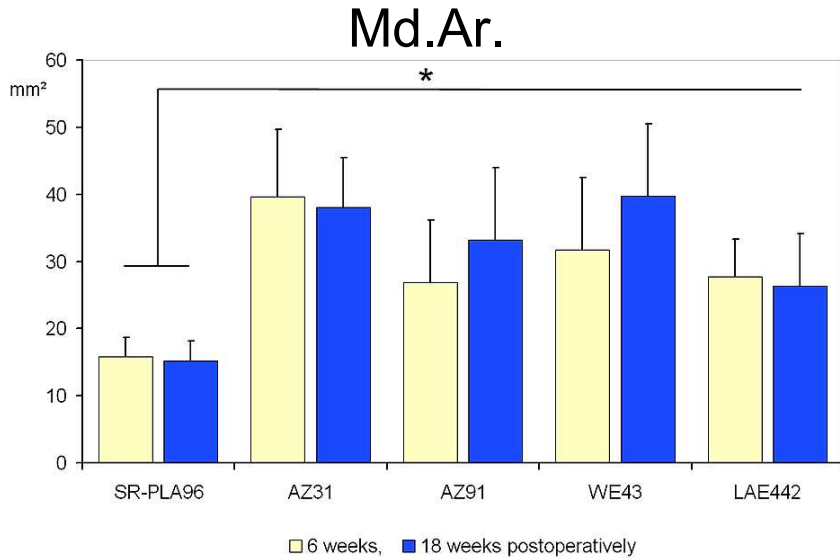
I = Implantat residual; E = endostal new bone formation;  
P = periosteal new bone formation; Bar = 1.5 mm

Witte et al., Biomaterials 26, 2005





# Histomorphometry of Mg-implants



## B.Ar./T.Ar

Table 1  
Bone area per total area (B.Ar./T.Ar) of implanted magnesium alloys and degradable polymer (SR-PLA96) in guinea pig femura at 6 and 18 weeks postoperatively

	6 weeks p-value	18 weeks p-value
AZ31/91	1.000	0.578
WE43/AZ31	0.847	0.806
WE43/AZ91	0.924	0.995
WE43/LAE442	0.646	1.000
LAE442/AZ31	0.996	0.730
LAE442/AZ91	0.980	0.999
SR-PLA96/AZ31	0.909	0.104
SR-PLA96/AZ91	0.963	0.828
SR-PLA96/WE43	1.000	0.605
SR-PLA96/LAE442	0.738	0.690

No significant differences were found between groups.

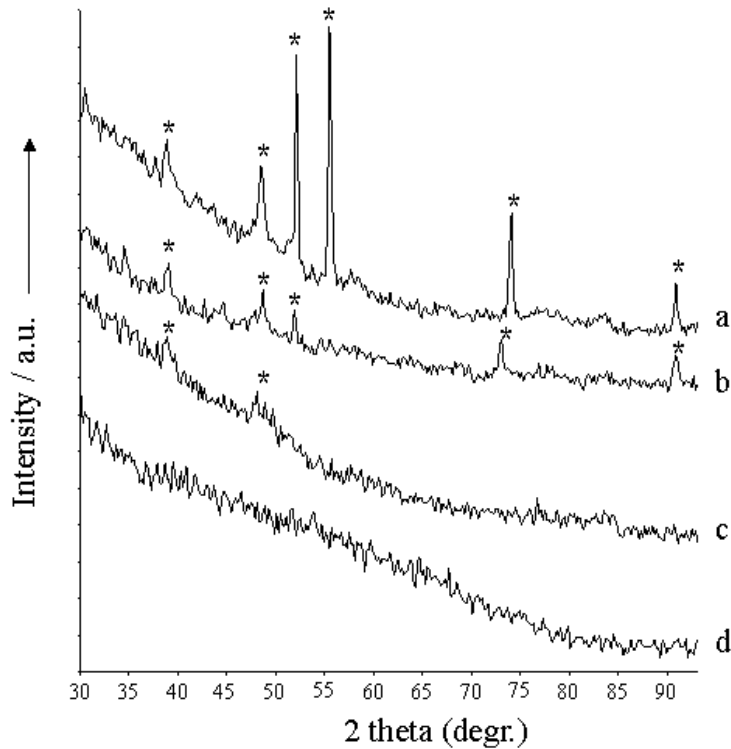
Physiological bone formation is enhanced by magnesium implants.

Witte et al., Biomaterials 26, 2005

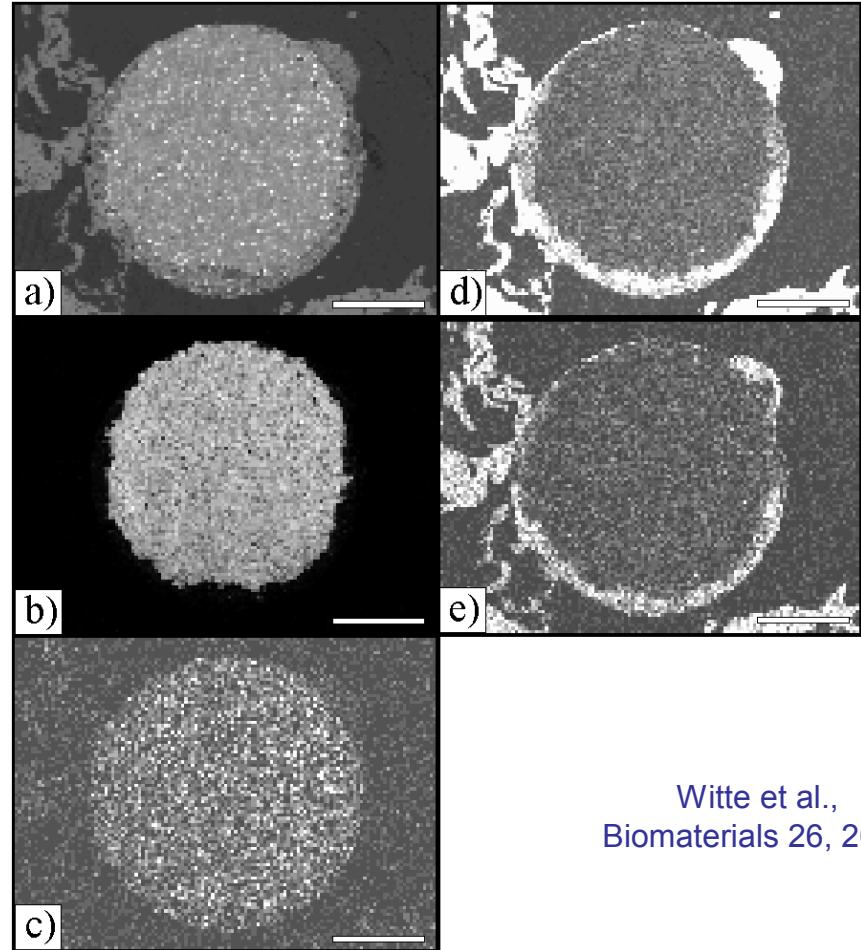




# Corrosion layer in-vivo



- a) magnesium residual
- b) corrosion layer
- c) bone
- d) embedding media



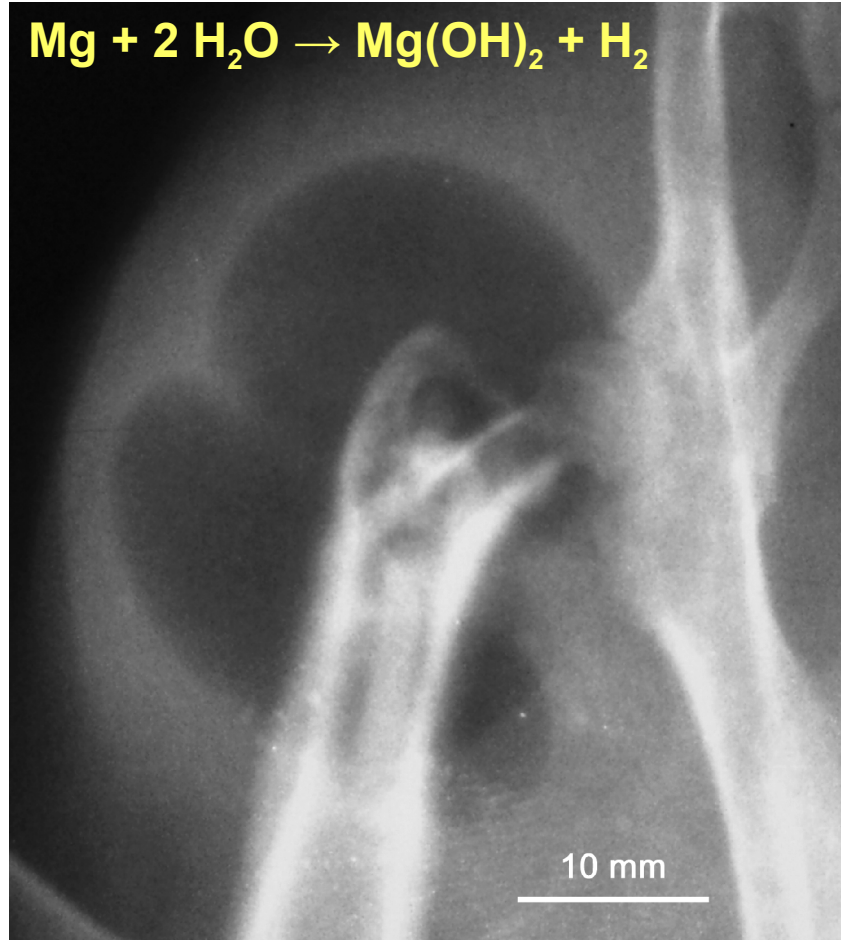
- a) BSC; b) magnesium; c) neodymium;
- d) calcium; e) phosphorus

Witte et al.,  
Biomaterials 26, 2005





# Subcutaneous gas cavities



1 g magnesium  $\gg$  1.08 l hydrogen gas

Witte et al., Biomaterials 26, 2005







# MMC - Novel biodegradable metals



Available online at [www.sciencedirect.com](http://www.sciencedirect.com)



Biomaterials 28 (2007) 2163–2174

**Biomaterials**

[www.elsevier.com/locate/biomaterials](http://www.elsevier.com/locate/biomaterials)

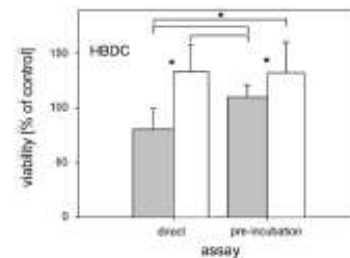
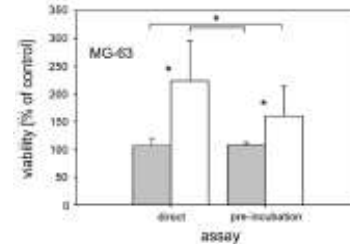
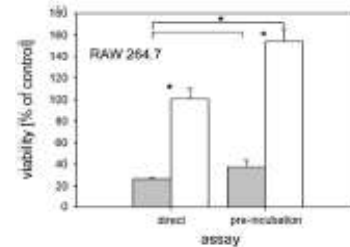
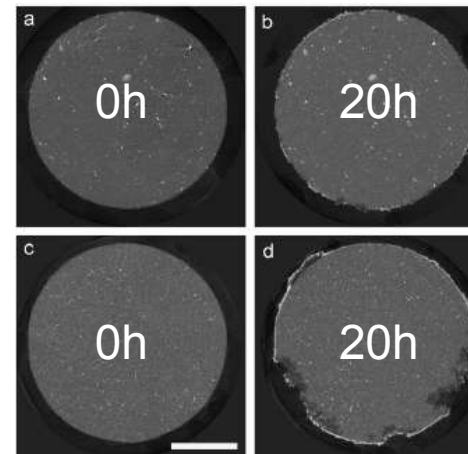
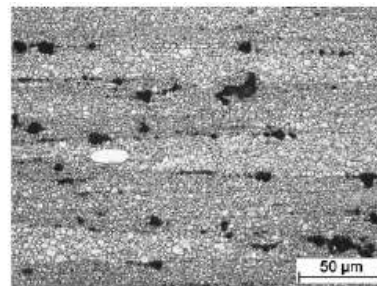
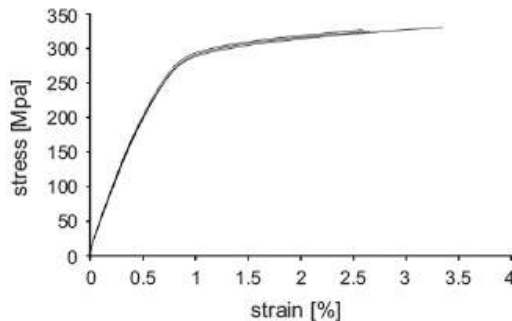
## Biodegradable magnesium–hydroxyapatite metal matrix composites

Frank Witte<sup>a,\*</sup>, Frank Feyerabend<sup>b</sup>, Petra Maier<sup>b</sup>, Jens Fischer<sup>a</sup>, Michael Störmer<sup>b</sup>,  
Carsten Blawert<sup>b</sup>, Wolfgang Dietzel<sup>b</sup>, Norbert Hort<sup>b</sup>

<sup>a</sup>Laboratory for Biomechanics and Biomaterials, Department of Orthopedic Surgery, Hannover Medical School,  
Anna-von-Borries-Str. 1-7, 30625 Hannover, Germany

<sup>b</sup>Institute of Materials Research, Magnesium Innovation Centre, GKSS Research Centre Geesthacht, Max-Planck-Str. 1, 21502 Geesthacht, Germany

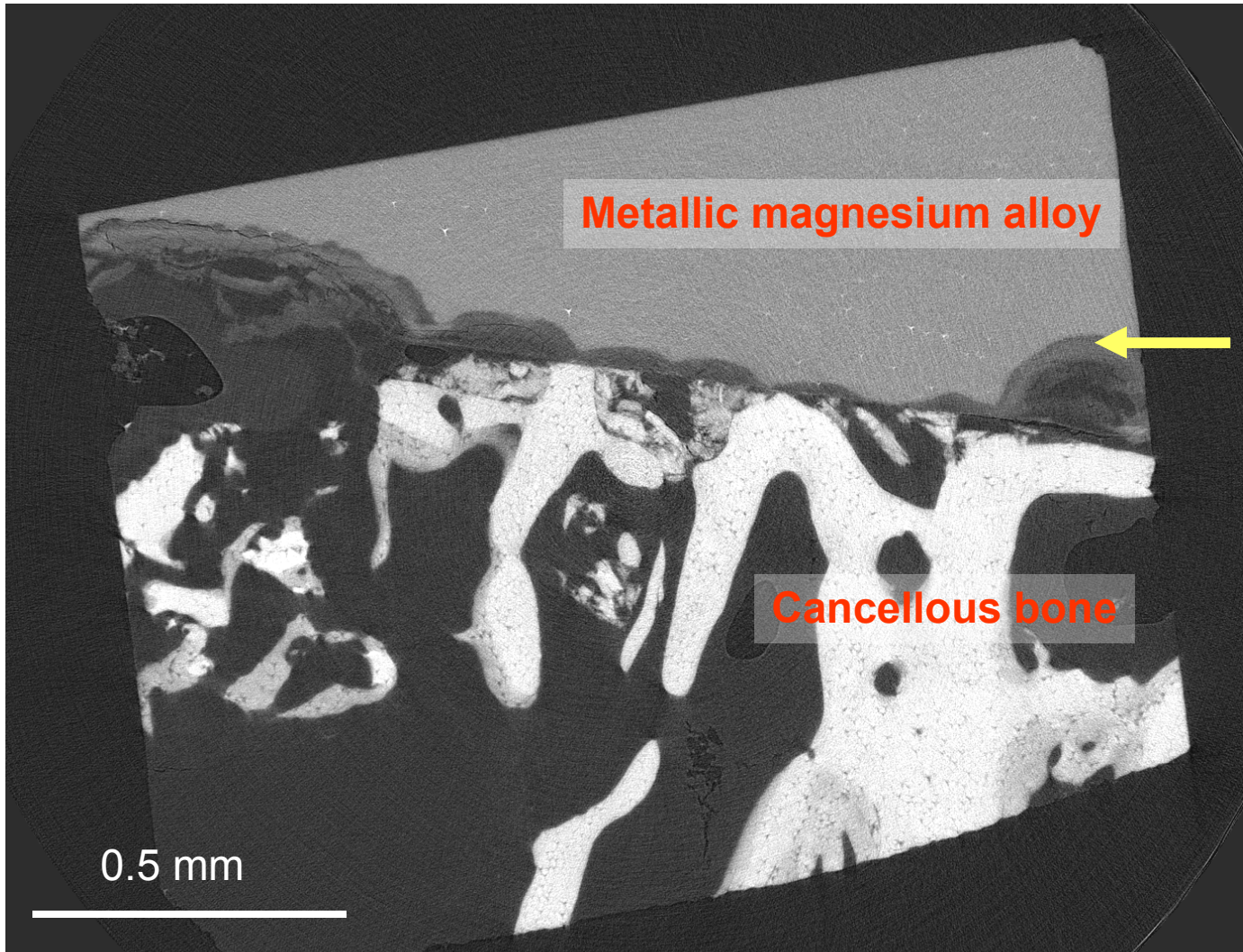
Received 7 August 2006; accepted 31 December 2006  
Available online 5 January 2007







# SR $\mu$ CT in attenuation mode



**Corrosion layer**

HASYLAB at DESY

beamline HARWI (W2)

photon energy: 31 keV

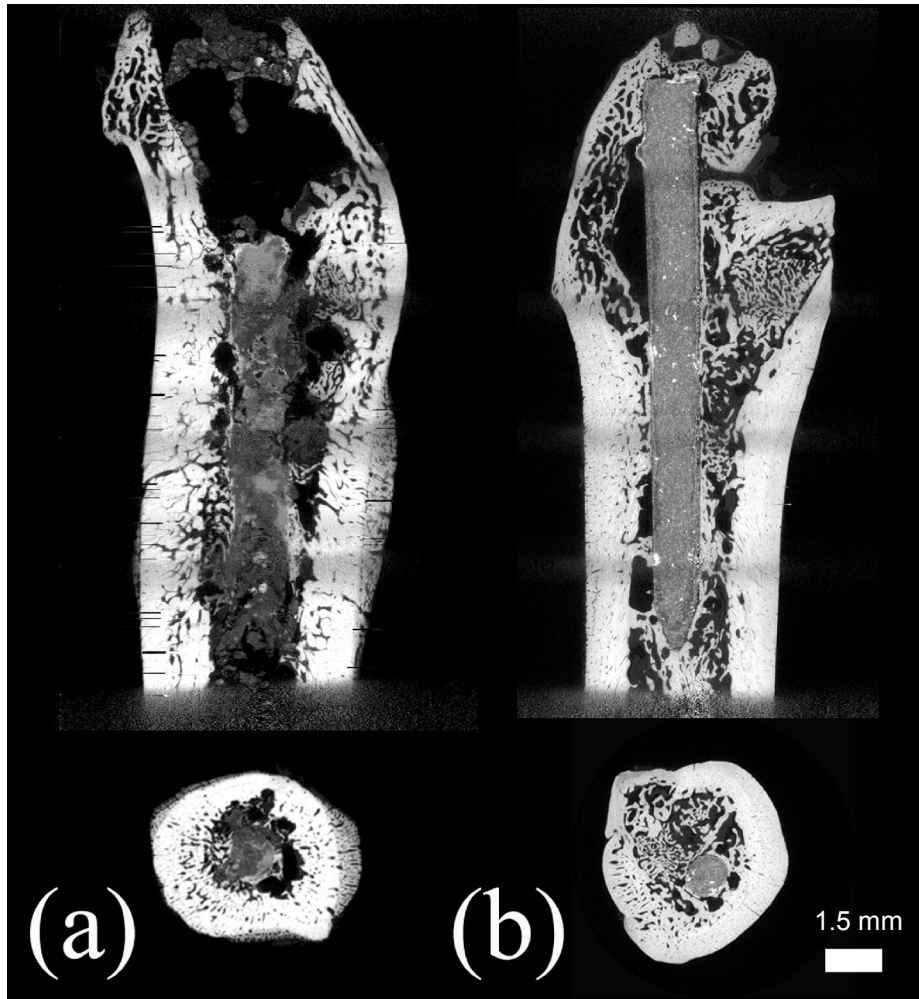
angular range 0-180°

stepwise 0.25°





# Analysis of the total specimens



AZ91D

LAE442

HASYLAB at DESY  
beamline HARWI (W2)  
photon energy: 31 keV  
angular range 0-180°  
stepwise 0.25°  
voxel edge length: 10  $\mu\text{m}$   
stack of 5 datasets

Witte et al,  
Biomaterials 27, 2006





# In vivo corrosion rate by SR $\mu$ CT

$$CR = \frac{W}{A \cdot t \cdot \rho} \quad (\text{Eq. 1})$$

Reference: ASTM G31-72, 2004

$$CR = \frac{\Delta V}{A \cdot t} \quad (\text{Eq. 2})$$

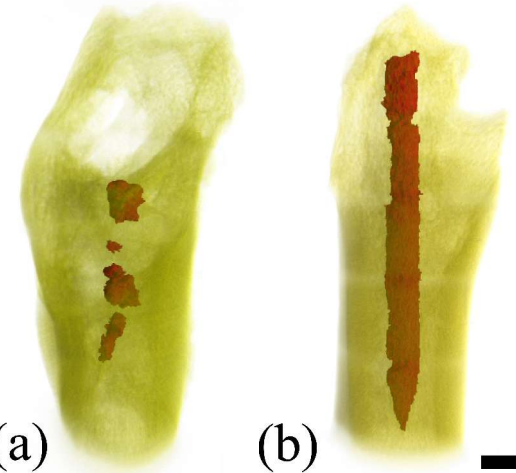
$CR$  is the corrosion rate

$W$  is the weight loss

$A$  is the original surface area exposed to the corrosive media

$t$  is the time of immersion

$\rho$  is the standard density



Witte et al., Biomaterials 27, 2006

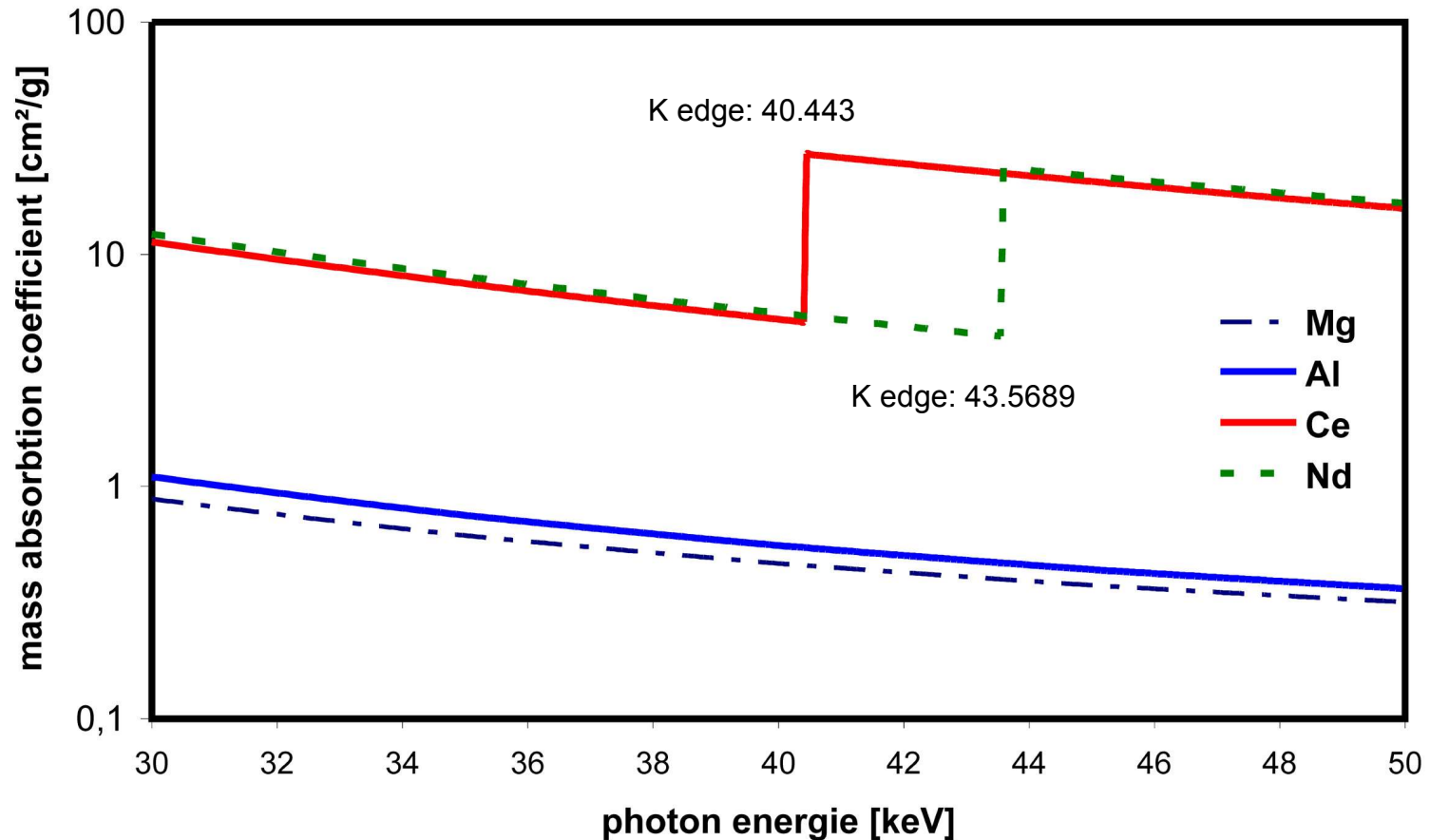






# Element-specific SR $\mu$ CT

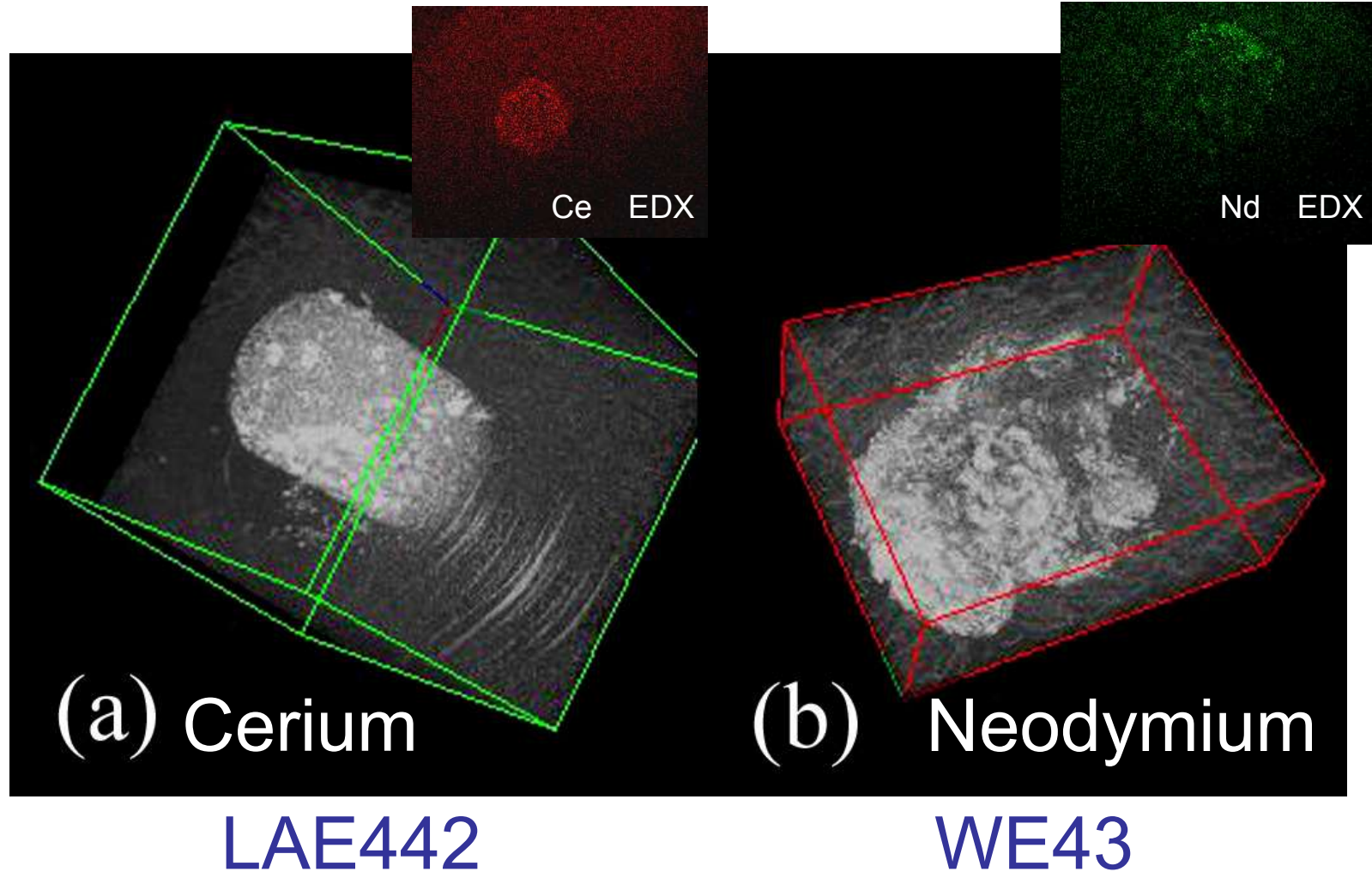
mass absorption coefficient vs. photon energy





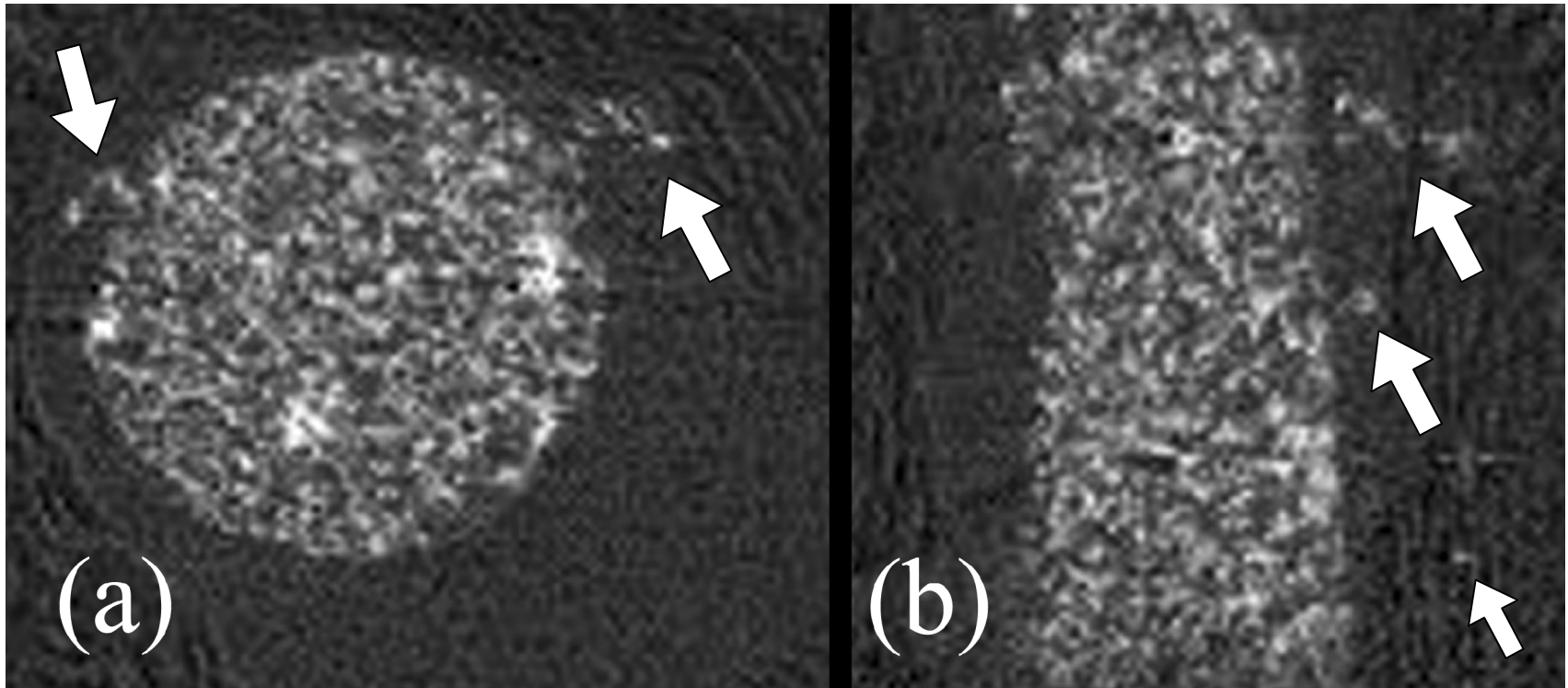


# Element-specific SR $\mu$ CT





# Spatial distribution of cerium (LAE442)





# Summary

## SR $\mu$ CT:

- became a great imaging tool
- has potential for quantitative analysis
- for structural biology and
- for interface analysis of biomaterials.





# Thank you very much !!!

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## Grants:

German Research Society, Colloabarative Research Project, SFB 599

Bundesministerium für Bildung und Forschung, BMBF

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