

Biomedical perspectives for spider silk fibres

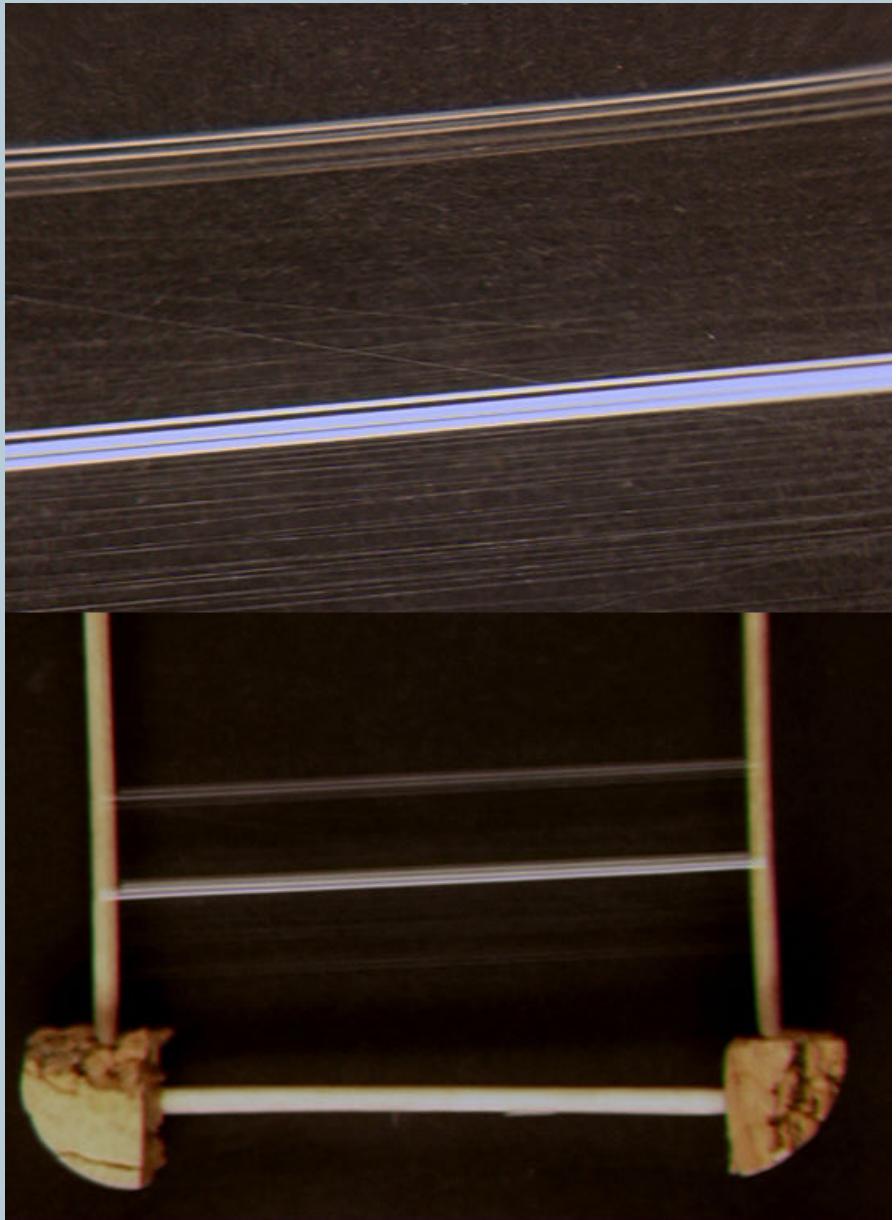
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Why spider silk?

- Mechanical properties, proteinic nature
- Different silk threads => properties => applications
 - Spider drag line
 - Spider cocoon silk
 - Synthetic spider silk

Silk: Spider dragline silk

- *Araneus diadematus* grown in the lab:
- The spider is fixated and dragline can be reeled off (1 rps/10min.): 1 spider \sim 100m = \sim 1mg
 - fine: D: 3-5 μm
 - strong/ elastic



Silk: Spider cocoon silk

- *Araneus diadematus* grown in the lab:
 - 1 spider => 1 cocoon => 10 mg of silk
- Easier to obtain than dragline
 - Very elastic
 - less strong than dragline



Synthetic spider silk

Genetic engineering of spider silk proteins

- Dupont: in Bacteria
- Hebrew University/Vollrath: Insect cells
- Nexia: in goats
- *Gatersleben*: in tobacco and in potato
- Still facing the problem of spinning the spider silk proteins.

Biomedical applications:

requirements

- Mechanical properties: strength/elasticity
- Sterilisable:
- Biocompatible:
- Biodegradable:
- Cell Support –not cytotoxic attachment/migration/expression/growth/differentiation/ ...
- ...

Mechanical properties

	Density [g cm ⁻³]	Tenacity [GPa]	Strain break [%]	Toughness [MJ m ⁻³]
nylon 6,6	1.1	0.95	18	80
kevlar 49	1.4	3.6	3	50
dragline of <i>A. diadema tus</i>	1.3	1.1	27	160
cocoon of <i>A. Diadema tus</i>	1,3	0,3	25-50	70
silk of the moth <i>B. mori</i>	1.3	0.6	18	70
wool	1.3	0.2	50	60
PLA	1.24	0.7	22	90
high-tensile steel	7.8	1.5	1	6

Elongation-strength curves cocoon cocoon silk

Elongation (%)	Strength (CN/dtex)	Series
0	0	diadematus
2	4.5	diadematus
5	0	diadematus
0	0	quadriatus
2	4.0	quadriatus
5	0	quadriatus
0	0	marmoratus
2	3.5	marmoratus
5	0	marmoratus

Elongation-strength curves dragline silk

Elongation (%)	Strength (CN/dtex)	Series
0	0	diadematus
2	8.5	diadematus
5	0	diadematus
0	0	quadriatus
2	8.0	quadriatus
5	0	quadriatus
0	0	marmoratus
2	7.5	marmoratus
5	0	marmoratus

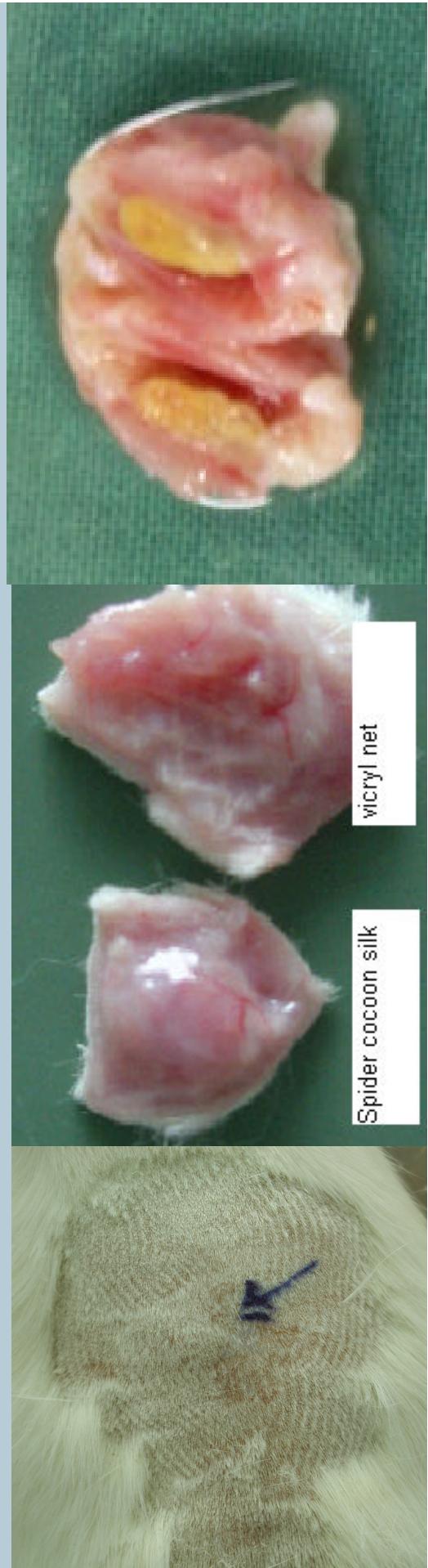
Sterilisation

- Ethylenoxide: toxic residues are left
- UV-sterilisation: makes the silk fibre less elastic, less strong and stiffer
- Steam sterilisation: same but less effect than UV-sterilisation and more effective against bacteria.

Conclusion: steam-sterillisation will be used

biocompatibility?

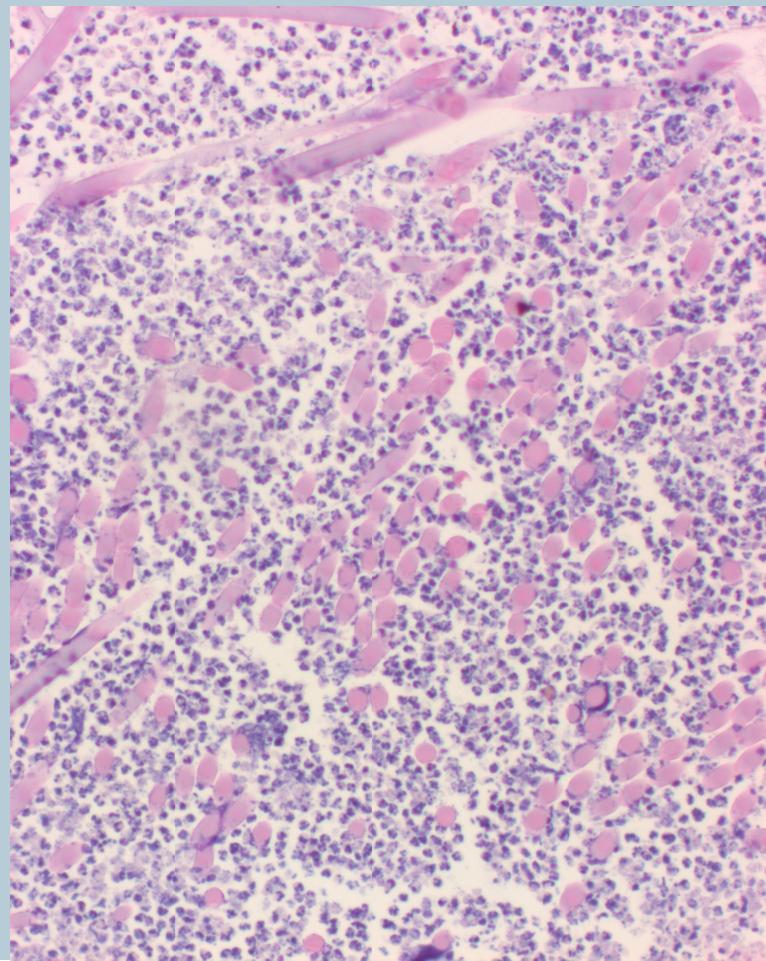
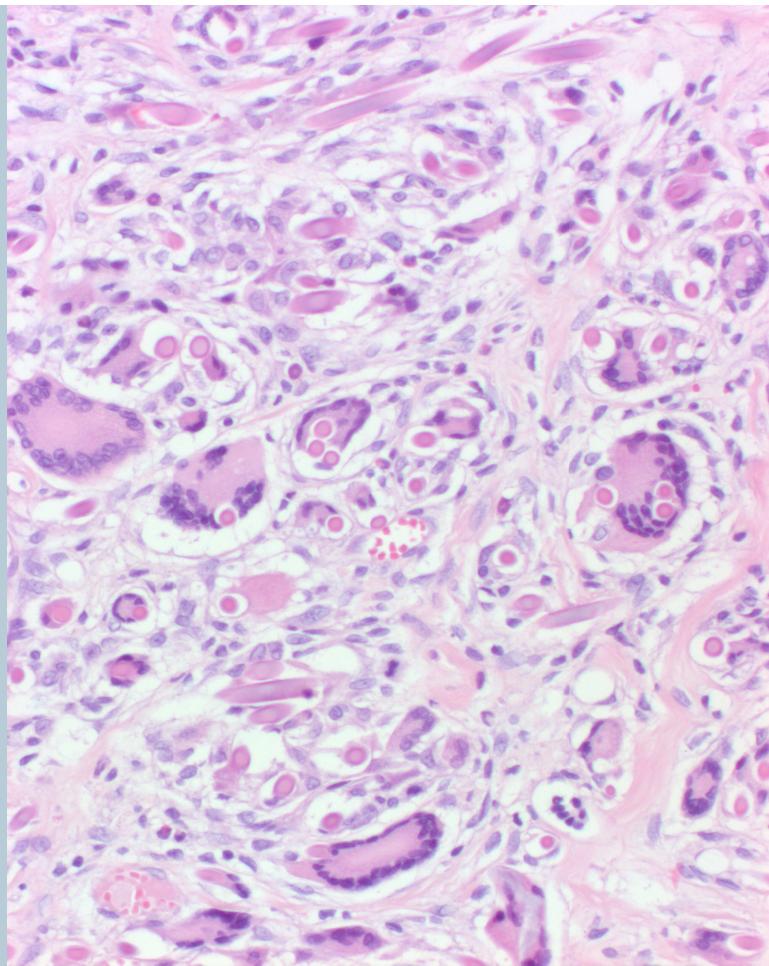
- **Silkworm silk:** fibroin with sericin-layer
 - problems: sericin shows biocompatibility problems
 - Can be removed by boiling in NaCO₃-solution
 - Silk fibroin is biocompatible
- **Spider silk:** *in vivo tests*
 - Spider silk has no sericin
 - Macroscopically: pain? Redness? Swelling? Heat?



biocompatibility

No enzymatic treatment: just autoclavage:

- more immune-cells



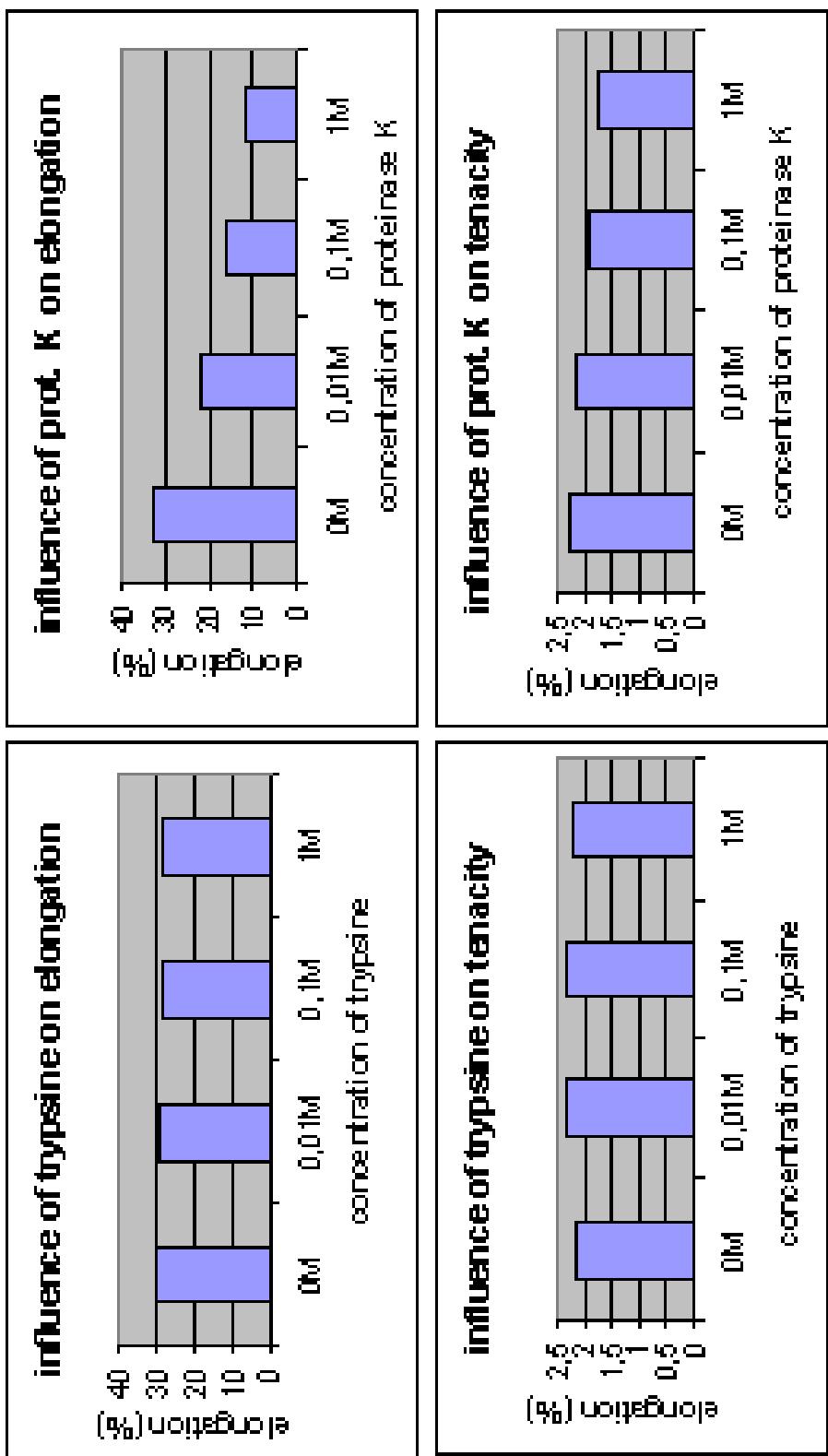
enzymatic treatment and autoclavage:

- less immune-cells

-Giant cells = typical towards inert material

-Fibrosis = recovery

Influence of enzymatic treatment



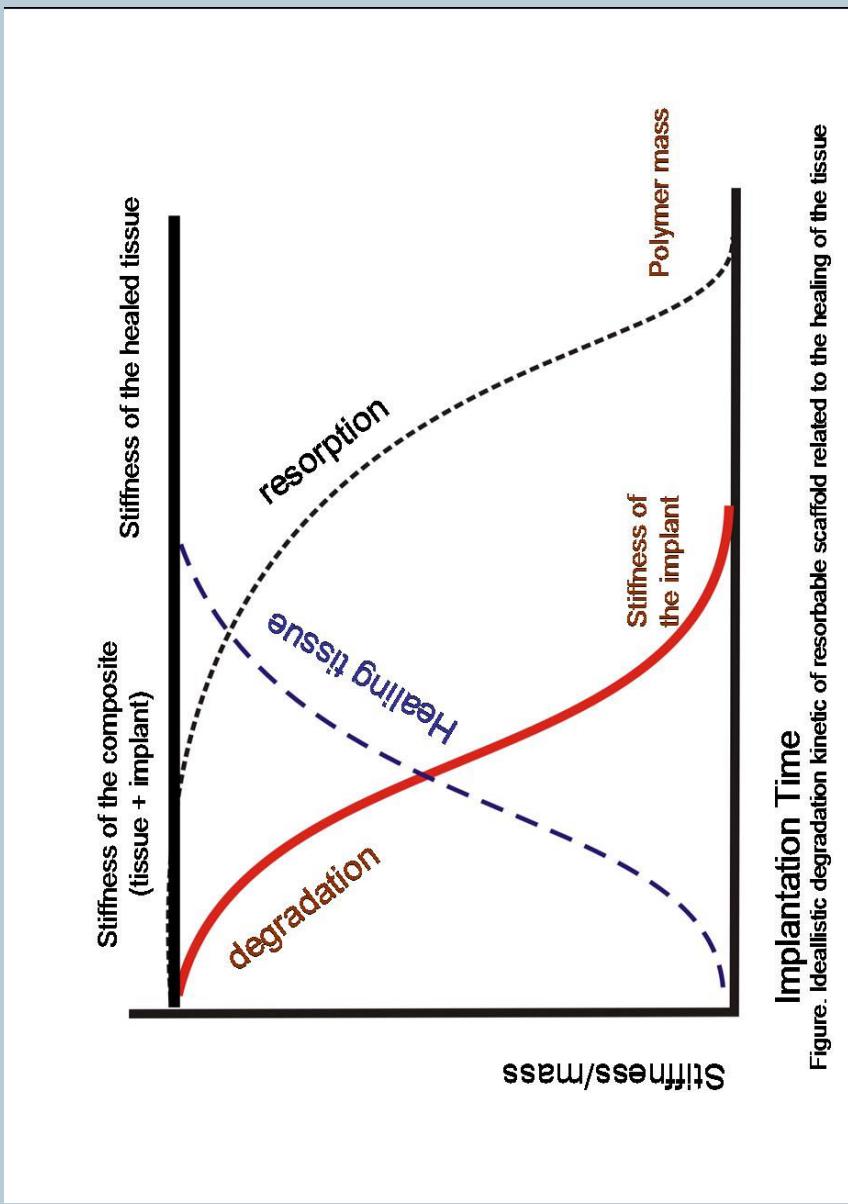
degradability

- In vivo: after 3 months silk is still there

Very slow degradability:

Negative for TE?

Speed of biodegradation
< speed of healing of the
tissue = good



Implantation Time
Figure. Idealistic degradation kinetic of resorbable scaffold related to the healing of the tissue

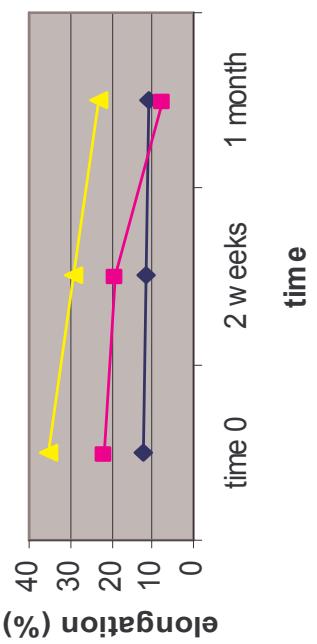
degradability

- In vitro: PBS, 37°C,, no stress, no enzymes

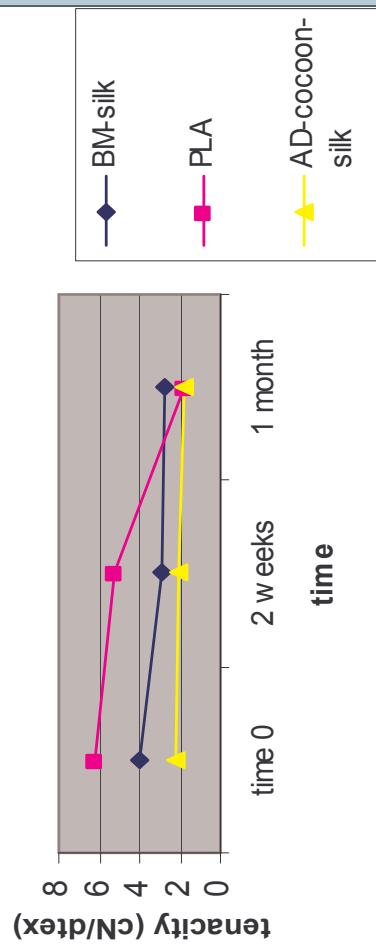
<u>elongation(%)</u>	BM-silk	PLA	AD-cocoon-silk
time 0	12,53	21,65	35,46
2 weeks	11,62	18,94	29,09
1 month	10,86	7,74	23,16

<u>tenacity (cN/dtex)</u>	BM-silk	PLA	AD-cocoon-silk
time 0	4,02	6,27	2,24
2 weeks	2,87	5,19	2,00
1 month	2,70	1,77	1,73

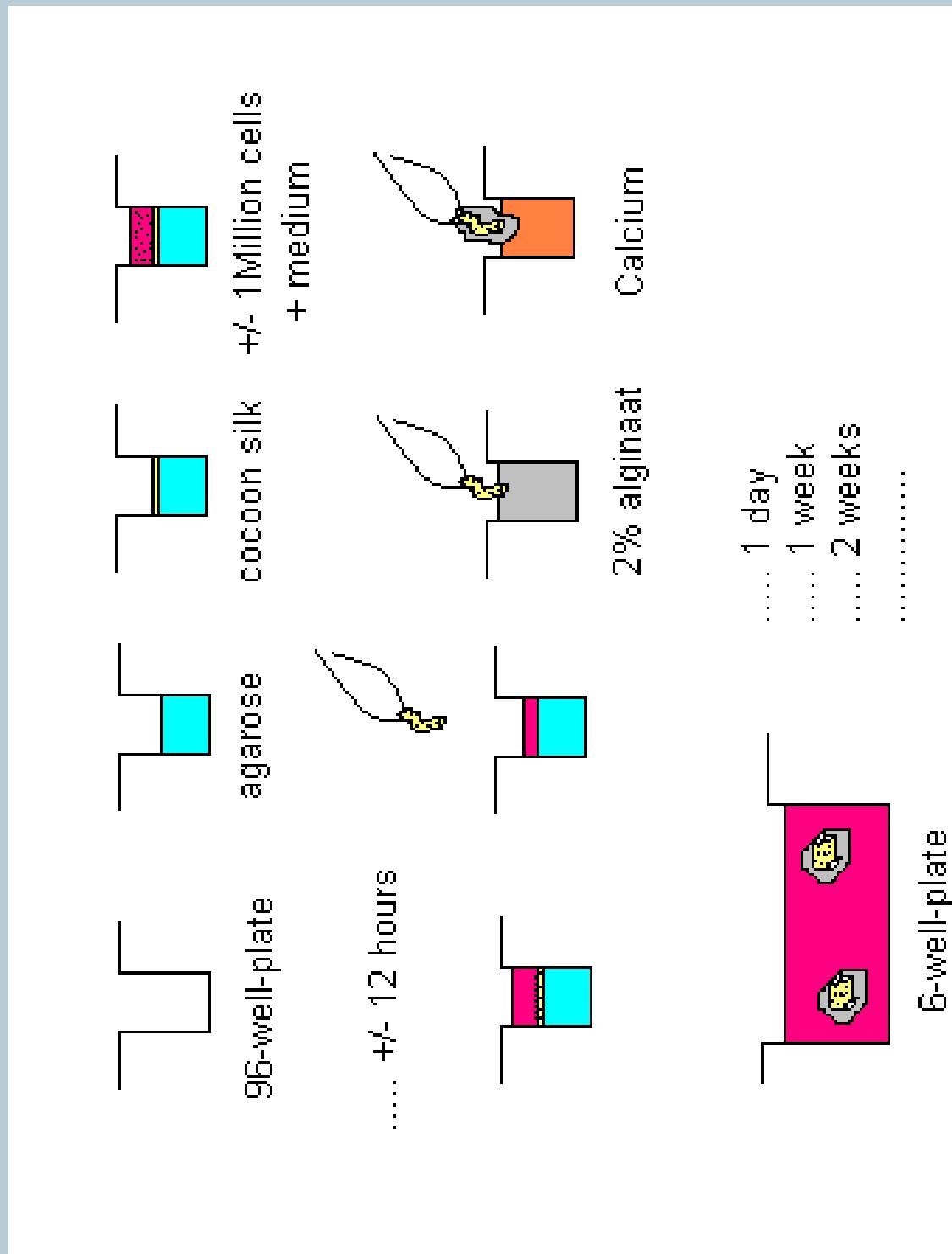
degradation of elongation



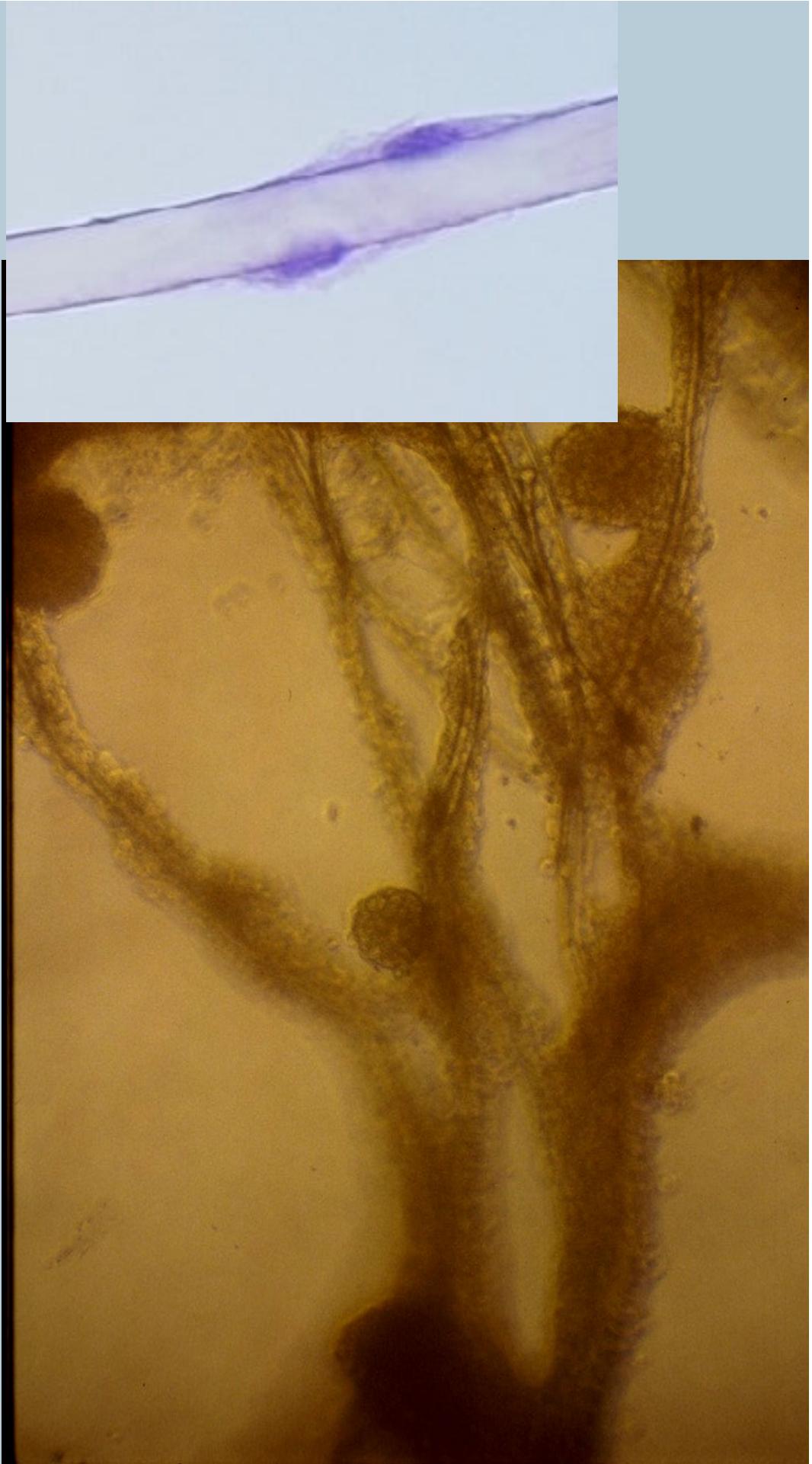
biodegradation of tenacity



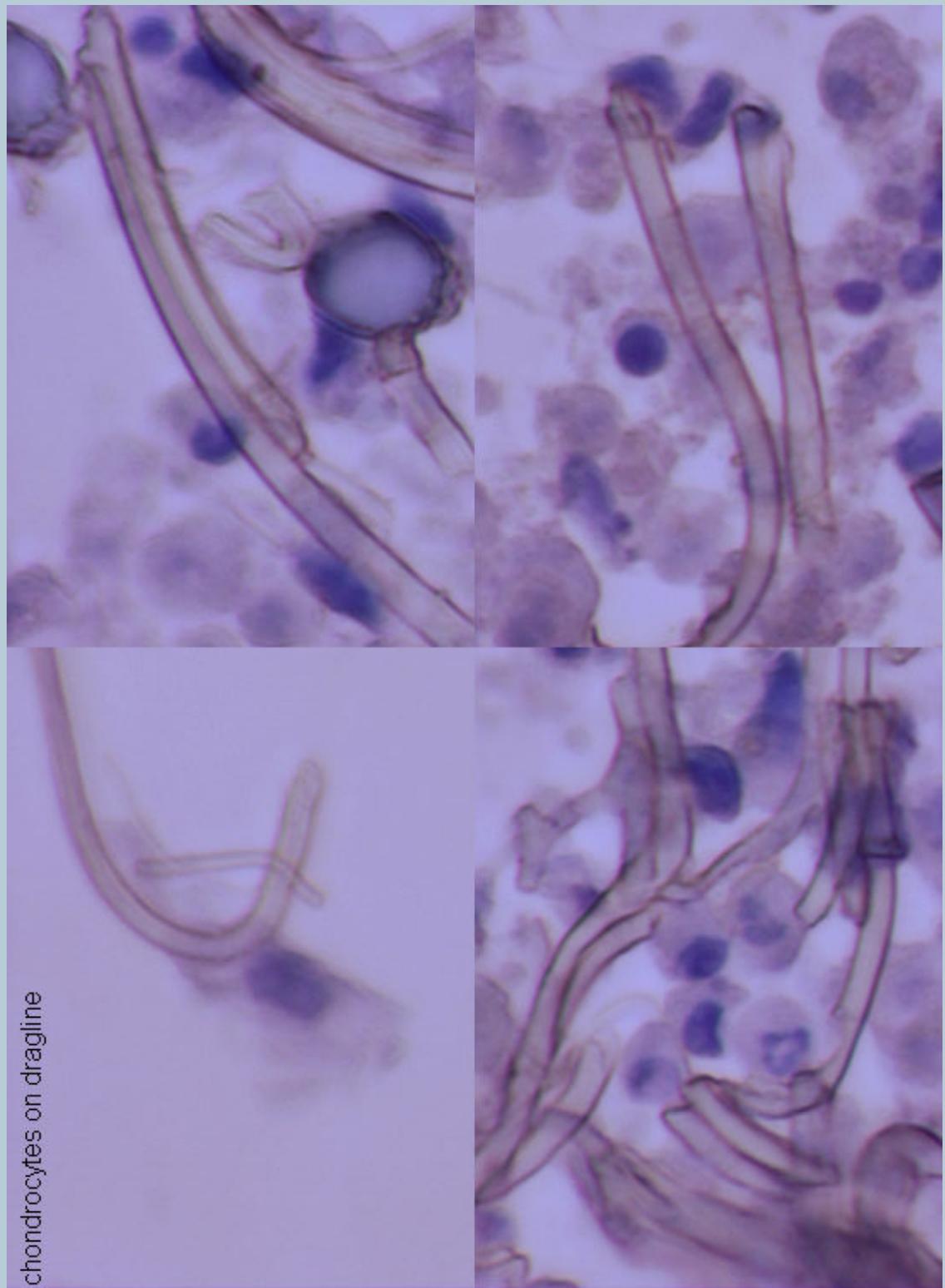
Cyto-intoxicity and cell-attachment



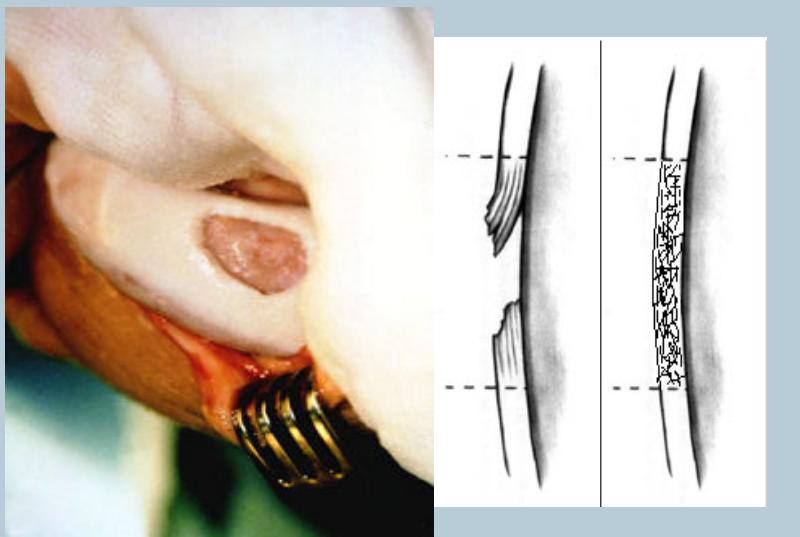
Cell-growth on spider cocoon silk fibres



Cell-growth on spider dragline fibres



Possible biomedical applications?



- Suture material.
- Wound dressing?
- Guiding of nerve cell or vascular tissue
- Scaffolds for
 - cartilage regeneration?
 - tendon repair?

Conclusion

- Biocompatible, slowly biodegradable, non-cytotoxic, sterilisable
- Natural and synthetical spider silk
- Usable in many applications

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