

# Biosensors with build-in logic for medical applications

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**Abstract.** The abstract should briefly summarize the contents of the paper in 150–250 words.

general concept

multisignal digital biosensors processing complex pattern of different physiological markers

practical considerations

challenges

**Keywords:** First keyword · Second keyword · Another keyword.

## 1 Introduction

overview of the novel research paradigm of digitally operating biosensors logically processing multiple biochemical signals through Boolean logic networks composed of biomolecular systems

### 1.1 Use of biosensors

### 1.2 innovation

### 1.3 how the technique works

## 2 Chancen

- biomedical monitoring(example, closed-loop- $\hat{c}$  patient tailored timely therapy, personalized medicine= sensing devices + delivery devices)
  - closed-loop - $\hat{c}$  patient tailored timely therapy possible
  - sensing devices + delivery devices = personalized medicine
  - example feedback-loop: diabetes: electrochemical glucose sensing element + insulin-delivery feedback loop
  - fast delivery in emergencies
- environmental monitoring
- national defense
- food safety

### 3 concept

- unconventional computing = quelle
- biocomputing multiple biochemical coupling of signal processing with chemical actuators

#### 3.1 Biocomputing

Definition

- single logic gates (mimicking Boolean) to small logic networks
  - biomolecular systems for processing chemical information
  - more complex than nonbiological systems
  - coupling enzymatic reactions (logic gates) with electronic transducers and signal-responding materials
- = enzyme logic gates

#### 3.2 Examples

theoretical

- glucose oxidase and catalase operating as logic gates:
- input : H<sub>2</sub>O<sub>2</sub> and glucose
- gluconic acid = biocatalytic oxidation of glucose
- only when both present optical output signal. = AND
- define logic values: small changes = 0 and large absorbance changes as 1 = AND
- similar possible with XOR, AND, OR, NOR, INHIBIT
- with logic gates with modular structure that enables their assembly in networks NAND/ NOR possible
- logic gates and their networks = biomolecular information processing systems
- = biosensoric systems with logically processed signals represented by various biomarkers (characteristic of different abnormal physiological conditions)

example pH

- pH changes in solution as logic responses to input signals
- AND invertase + glucose oxidase (from 5.8 to 3.5)
- OR esterase and glucose oxidase in glucose and ethyl butyrate - when one of both present - acidification
- neutral pH = 3.5

Conclusion:

- don't solve real computing problem nor operate as useful biosensors
- represent first step toward the development of digital biosensors
- funfact optimization of enzymatic reaction, up to 10 logic gates concatenated with low noise in the system

### ENzyme logic system recognizing various injury-related physiological conditions

- types of injuries result in concentrations of chemical substances in the body
- example: lactate oxidase, horseradish peroxidase and glucose dehydrogenase  
= designed to process biochemical information related to pathophysiological conditions from brain injury
- markers: glucose(hemorrhagic shock), lactate(rhagic shock or traumatic brain injury) and norepinephrine(traumatic injury)
- logic 0 = normal concentrations
- change results into different numbers 1,2,3 - convenient
- = biocomputing logic system
- challenge: difference between normal and unnormal minimal =, not linear, should be sigmoidal

## 4 challenges

prospects fundamental and practical challenges

- attention to composition, preparation and immobilization of the biocomputing surface layer
- success depends in part on immobilization of the biocomputing reagent layer
- system scalability
- efficient transduction of the output signals
- high fidelity

stuff missing

da war was mit wie viel können hintereinander geschaltet werden

### 4.1 Enzyme logic circuits-scaling up the system complexity

- main challenge: scaling up the complexity of the systems by networking the individual parts of a logic circuit
- addressed experimentally when designing networks composed of concatenated enzyme logic gates
- assembled logic networks analyzed theoretically for optimization and noise reduction; coupling output signals with electronic transducers and bioelectronic devices

### 4.2 Biomolecular logic gates designed for biomedical analytical applications

problems not addressed yet in biomolecular information processing systems:

- logic 0 values were defined as the absence of the enzymes
- logic 1 not always correspond to the concentration expected in vivo /not normal physiological concentrations
- input not justified to their biomedical meaning

## 5 advantages

- multiple target analytes(inputs) (enzymgates)/biochemical inputs
- high-fidelity biosensing compared
- rapid and reliable assessment of physiological condition (enzymes + automatically processing)
- optimal timely therapeutic intervention
- realization of closed-loop systems (sense/act/treat)
- missing example fields like diabetes (siehe text)

## 6 Conclusion

## 7 Idee des Aufbaus

- Abstract: Biosensoren, logic gates, possibilities, challenges and future work
- introduction Definition, possible fields(examples), special,... loops
- The idea Theory and practical
- Two examples
- Layers, challenges and future work
- conclusion

## References

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3. Joseph Wang, Evgeny Katz, Digital biosensors with build-in logic for biomedical applications- biosensors based on a biocomputing concept, in: Anal Bioanal Chem(2010) 1591-1603