

# Ion-Selective Field Effect Transistors (ISFETs)

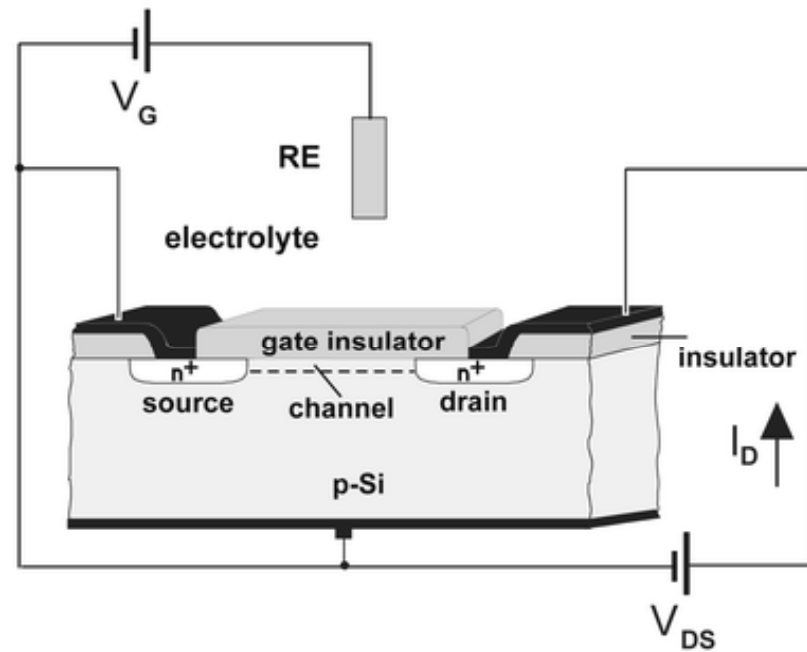


Common transducer in commercial pH meters.

Gate insulator materials:  
 $\text{Si}_3\text{N}_4$ ,  $\text{Al}_2\text{O}_3$ ,  $\text{Ta}_2\text{O}_5$

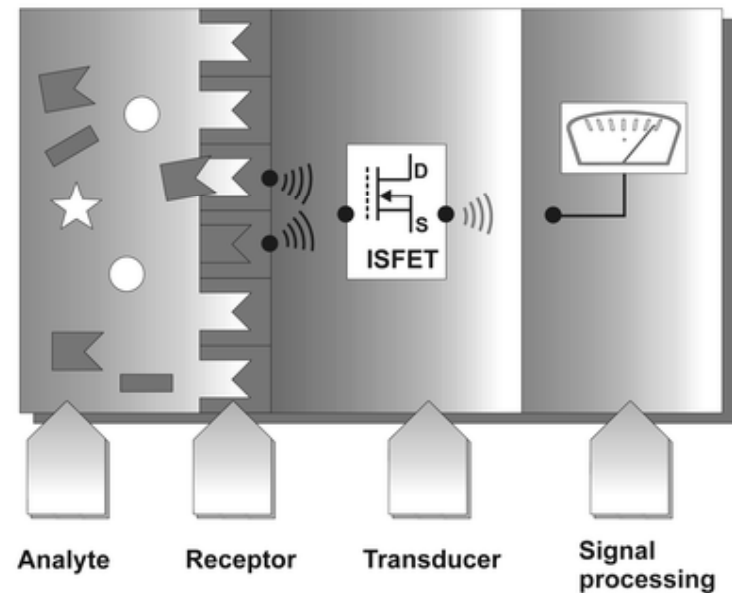
Surface of gate insulator becomes hydrolyzed in aqueous solution.

-OH groups on surface experience varying degrees protonation depending on the surrounding pH



# BioFET

What about BioFETs?



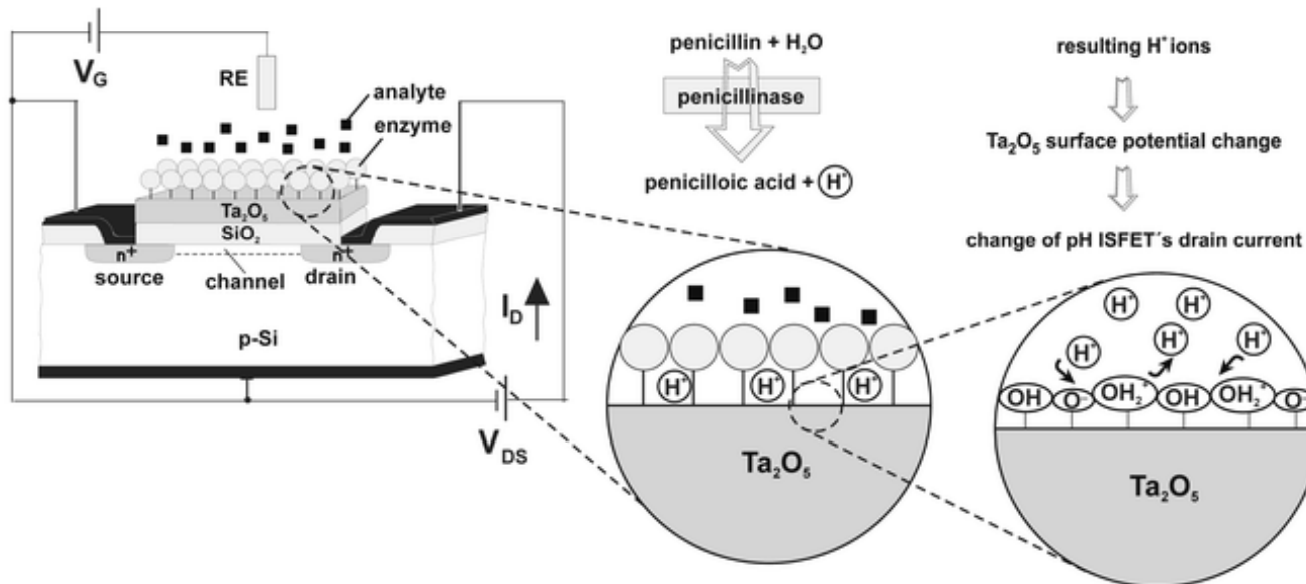
- Sensitivity to any electrical interaction nearby the gate insulator can be extended to biosystems.
- Any biochemical reaction that leads to chemical or electrical changes at this interface can be measured.

 ISFET coupled with a bioreceptor

1. Potential changes caused by a catalytic reaction product  
*between enzyme and substrate*
2. Potential changes caused by surface polarization effects  
or through change in dipole moments  
*antigen-antibody affinity reactions*  
*DNA hybridization*
3. Potential changes caused by biological processes as a  
results of more sophisticated (bio)chemical processes  
*action potential of nerve cells*

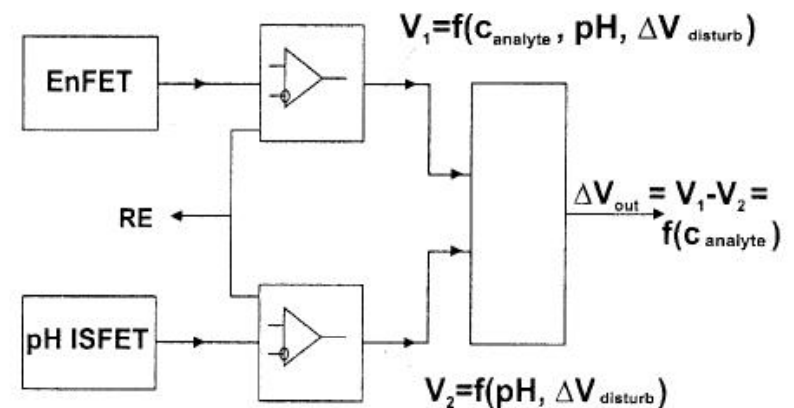
## Example: Enzyme -modified FET (EnFET)

- Enzyme used as bioreceptor due to specific binding
- Enzymes are bound to gate insulator through a number of ways:
  - physical or chemical adsorption
  - entrapment within polymeric matrices
  - cross-linking through cross-linking agents
- General working principle: during the enzymatic reaction with the substrate, products are generated which change the local solution concentration. This change is monitored by the ISFET



**Table 2** Recently developed EnFETs including the enzyme system used and the analyte to be detected

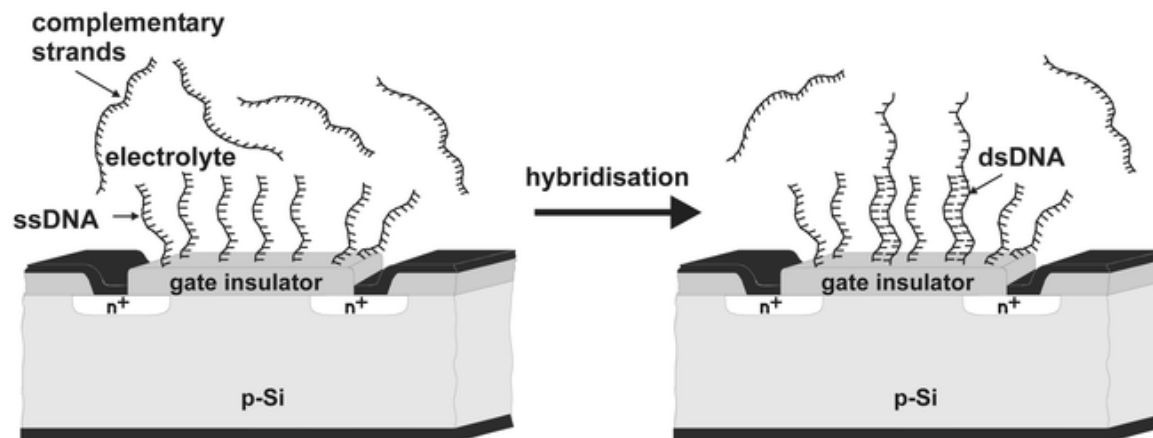
Analyte	Enzyme system
Glucose	Glucose oxidase <sup>90-98,100,129,130</sup> Glucose oxidase/MnO <sub>2</sub> powder <sup>99</sup> Glucose dehydrogenase <sup>101</sup>
Urea	Urease <sup>87,95,101-108</sup> Penicillin penicillinase <sup>109-113,115</sup> Penicillin G acylase <sup>114</sup>
Sucrose	Invertase/mutarotase/glucose oxidase <sup>1</sup> Invertase/glucose dehydrogenase <sup>101,1</sup>
Maltose	Maltase/glucose dehydrogenase <sup>101</sup>
Ethanol	Alcoholdehydrogenase/aldehyde dehydrogenase <sup>101</sup>
Lactose	β-Galactosidase/glucose dehydrogenase <sup>116</sup> β-Galactosidase/galactose dehydrogenase <sup>101</sup>
Ascorbic acid	Peroxidase <sup>98,118</sup>
Creatinine	Creatinine deiminase <sup>126</sup>
Formaldehyde	Alcohol oxidase <sup>124,125</sup>
Acetylcholine	Acetylcholinesterase <sup>95,119</sup>
Organophosphate compound (paraoxon)	Organophosphate hydrolase <sup>122</sup>
Fluorine-containing organophosphates	Organophosphorus acid anhydrolase <sup>123</sup>



**Fig. 5** pH ISFET/EnFET differential arrangement. The pH ISFET acts as 'reference' system; it is built-up in the same way as the EnFET but without immobilised enzyme membrane.

## Example: Gene -modified FET (GenFET)

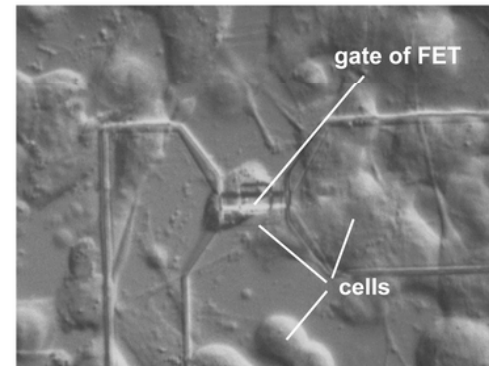
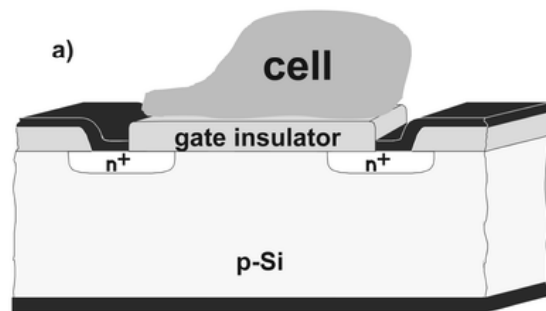
- Fast, simple and inexpensive analysis of nucleic acid samples
- Applications including diagnosis of genetic diseases, detection of infectious agents, drug screening, etc.



- Immobilize well-defined sequences of DNA onto transducer
- Specific recognition occurs between two DNA strands through hybridization
- Hybridization event (which induces a dipole  $\Delta$ ) converted into a measurable signal
  - ✓ Remains relatively unexplored as a DNA-based sensor
  - ✓ Counter ions can interfere with measurement

# Example: Cell-based BioFET

- Whole cells are used as the recognition element
- A single cell or cell system is coupled to the gate insulator of a ISFET
- Applications include the study of the effects of chemical and physical stimulus on the biological system.
- Examples include pharmaceutical compounds, toxic substances, pollutants, etc.



- Changes monitored by the ISFET include extracellular pH, ion concentration,  $O_2$  consumption,  $CO_2$  production, redox potential, other metabolic products, neuronal potentials, ...

