

## ES1

### Electrochemical Sensors for DNA Hybridization by Electron Transfer

전자전달을 이용한 DNA 검출의 전기화학적 센서

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Identifying infectious organisms, quantitating gene expression, and sequencing genomic DNA on chips all rely on the detection of nucleic acid hybridization. Described here is a novel assay for detection of the hybridization of products of the polymerase chain reaction using electron transfer from guanine to a transition-metal complex. The hybridization assay was modeled in solution by monitoring the cyclic voltammetry of  $\text{Ru}(\text{bpy})_3^{2+}$  ( $\text{bpy}=2,2'$ -bipyridine) in the presence of a probe strand containing only A, T, and C prior to and after hybridization to a complement that contained seven guanines, which led to high catalytic current due to the oxidation of guanine by  $\text{Ru}(\text{bpy})_3^{3+}$ . To allow recognition of all four bases in the target sequence, it was shown that inosine 5'-monophosphate was 3 orders of magnitude less reactive than guanosine 5'-monophosphate, suggesting that effective hybridization sensors could be realized by immobilization of probe strands in which inosine was substituted for guanosine; hybridization to guanosine-containing target strands would then provide high catalytic currents. A sensor design was tested in a model system for the detection of a synthetic 21-mer oligonucleotide patterned on the sequence of the ras oncogene, which gave an increase in charge collected of  $35 \pm 5 \mu\text{C}$  after hybridization and of only  $8 \pm 5 \mu\text{C}$  after exposure to noncomplementary DNA. Independent quantitation of probe and target by radiolabeling showed that the hybridized electrode contained  $3.0 \pm 0.3 \text{ ng}$  of target. New sensor electrodes were then prepared for the detection of PCR-amplified genomic DNA from herpes simplex virus type II, genomic DNA from *Clostridium perfringens*, and genomic RNA from human immunodeficiency virus and gave an additional charge of 35-65  $\mu\text{C}$  for hybridization to complementary amplicon and only 2-10  $\mu\text{C}$  after exposure to noncomplementary DNA.