36 MOLECULARLY IMPRINTED POLYMERS: UNIQUE POSSIBILITIES FOR ENVIRONMENTAL MONITORING

Ian A. Nicholls Karina Adbo Håkan S. Andersson Per Ola Andersson Jimmy Hedin Dahlström Jesper G. Karlsson Jenny P. Rosengren Johan Svenson Susanne Wikman Department of Chemistry & Biomedical Sciences, University of Kalmar SE-391 82 Kalmar, Sweden

ABSTRACT

Molecular imprinting provides a useful complement to the use of biomolecules for the development of sensors for the analysis of trace substances on account of their highly selective recognition characteristics. Moreover, the stabilities make them ideal for use in applications not normally suitable for the use of biomolecules. The nature of the technique and its use in a number of analysis formats suitable for use in environmental monitoring are presented.

KEYWORDS

Molecular imprinting, molecularly imprinted polymer, sensor, environmental analysis.

The need for improved sensitivity and robustness are two major factors motivating the development of new analysis techniques. Nature's capacity to produce highly selective recognition systems (*e.g.* antibody-antigen, receptor-ligand, enzyme-substrate) has made the use of biomolecules as recognition elements in analytical techniques, in particular for the determination of trace materials in complex samples, most rewarding. Nonetheless, the stability of biomolecules with respect to their operational tolerances (*e.g.* temperature, pH, ionic strength and organic solvents) is generally limited.

https://doi.org/10.15626/Eco-Tech.2001.035

KALMAR ECO-TECH'01 Leachate and Wastewater Treatment with High-Tech and Natural systems KALMAR, SWEDEN, November 26-28, 2001



Figure 1: A highly schematic representation of the molecular imprinting process. A monomer (or monomer) mixture with chemical functionality complementary to that of the template (T) is allowed to form solution adducts(2) through the complementarily interacting functionalities (reversible covalent or non-covalent interactions). Polymerisation in the presence of a cross linking agent (3), followed by removal of the template, leads to the defining of recognition sites of complementary steric and functional topography to the template molecule.

Molecular imprinting, Figure 1, is an emerging technique for producing highly selective synthetic receptors for an ever-increasing range of biochemical and chemical structures (1-4). The method involves the formation of cavities in a synthetic polymer matrix that are of complementary functional and structural character to a template molecule. The ability of molecularly imprinted polymers (MIPs) to selectively recognise and bind the template structure in the presence of closely related chemical species has led to them being used in a range of biomedical and biotechnological applications. As antibody combining site mimics, MIPs have demonstrated binding affinities and cross-reactivity profiles comparable to their biological counterparts and have even been employed as substitutes for biological antibodies in environmental analysis and for medical diagnostic assays. They have also been used as highly selective chiral chromatographic stationary phases. More recently, the first real attempts have been made to utilise the highly selective recognition of MIPs for producing enzyme mimics (5-7), following a strategy similar to that used for the production of catalytic antibodies. While molecular imprinting carries strong parallels to the exploitation of the immune system for producing antibodies selective for molecular features it also offers significant advantages, namely: no requirement for laboratory animals, time efficiency, relatively low cost of production, no hapten conjugation protocols and non-immunogenic

286 Ian A. Nicholls, Karina Adbo, Håkan S. Andersson, Per Ola Andersson, Jimmy Hedin Dahlström, Jesper G. Karlsson, Jenny P. Rosengren, Johan Svenson, Susanne Wikman, Sweden

KALMAR ECO-TECH'01 Leachate and Wastewater Treatment with High-Tech and Natural systems KALMAR, SWEDEN, November 26-28, 2001

substances pose no problem. Moreover, these materials are stable to extremes of pH, temperature and organic solvents (8), and offer significant advantages over the use of biomolecules in terms of the development of robust analytical systems, such as for implementation in the field.

The flexibility of the technique with respect to the choice of template, polymer composition and polymerisation format allows for their ready incorporation into a number of sensorformats. Furthermore, the resilience of these materials offers significant advantages over. MIPs have been utilized in a number of sensor formats for trace analysis. Optical detection has been the focus of much work, as exemplified by a fluoroimmuno-assay for the detection of 2,4-D and related substances (9) and a surface plasmon resonance method for the detection of drugs used in the treatment of asthma (10). MIP-based quartz crystal microbalances have been developed for the detection of 2-methylisoborneol, a substance responsible for the particular smell of stagnant water (11). A number of electrochemical sensors have been investigated, the most noteworthy being one for the detection of products resulting from the degradation of the nerve gas sarin (12). Competitive binding assays, for example for the detection of triazine herbicides (13), have also been developed.

Collectively, strong argument can be presented for the continued development of these MIPs, and their implementation in sensor development, especially for situations requiring robust analysis procedures such as in industrial and field applications.

REFERENCES

- 1. Sellergren, B. (2000). Imprinted polymers with memory for small molecules, proteins, or crystals. *Angew. Chem. Int. Ed.* 39, 1031-1037.
- 2. Andersson, H.S., Nicholls, I.A. (1997). Molecular imprinting: recent innovations in antibody and enzyme mimicking synthetic polymers. *Recent Res. Develop. Pure Appl. Chem. 1*, 133-157.
- Haupt, K., Mosbach, K. (1998). Plastic antibodies: developments & applications Trends Biotechnology 16, 468-475.
- Asanuma, H., Hishiya. T., Komiyama, M. (2000). Tailor-made receptors by molecular imprinting. Adv. Mater. (2000) 12, 1019-1030.
- Matsui, J., Nicholls, I.A., Karube, I., Mosbach, K. (1996). Carbon-carbon bond formation using substrate selective catalytic polymers prepared by molecular imprinting: an artificial class II aldolase. J. Org. Chem. 61:5414-5417.
- Strikovsky, A.G., Kasper, D., Grun, M., Green, B.S., Hradil, J., Wulff, G. (2000). Catalytic molecularly imprinted polymers using conventional bulk polymerization or suspension polymerization: Selective hydrolysis of diphenyl carbonate and diphenyl carbamate. J. Am. Chem. Soc. 122, 6295-6296.
- 7. Sellergren, B., Karmalkar, R.N., Shea, K.J. (2000). Enantioselective ester hydrolysis catalyzed by imprinted polymers. J. Org. Chem. 65, 4009-4027.
- Svenson, J., Nicholls, I.A. (2001). On the thermal and chemical stability of molecularly imprinted polymers. *Anal. Chim. Acta* (2001) 435, 19-24.
- Haupt, K., Mayes, A.G., Mosbach, K. (1998) Herbicide assay using an imprinted polymer based system analogous to competitive fluoroimmunoassays. Anal. Chem. *Anal. Chem.* 70, 3936-3939.

Ian A. Nicholls, Karina Adbo, Håkan S. Andersson, Per Ola Andersson, Jimmy Hedin Dahlström, 287 Jesper G. Karlsson, Jenny P. Rosengren, Johan Svenson, Susanne Wikman, Sweden

KALMAR ECO-TECH'01 Leachate and Wastewater Treatment with High-Tech and Natural systems KALMAR, SWEDEN, November 26-28, 2001

- Lai, E.P.C., Fafra, A., Vandernoot, V.A., Kono, M., Polsky, B. (1998). Surface Plasmon Resonance Sensors Using Molecularly Imprinted Polymers for Sorbent Assay of Theophylline, Caffeine, and Xanthine. *Can. J. Chem.* 76 265-273.
- Ji, H.-S., McNivan, S., Ikebukuro, K., Karube, I. (1999). Anal. Chim. Acta Selective Piezoelectric Odor Sensing Using Molecularly Imprinted Polymers. 390, 93-100.
- Jenkins, A.L., Uy, O.M., Murray, G.M. (1999). Anal. Chem. Based Lanthanide Luminescent Sensor for Detection of the Hydrolysis product of the nerve agent soman in water. 70, 373-378.
- Piletsky, S.A., Piletskaya, E.V., El'skaya, A., Levi, R., Yano, K., Karube, I. (1997). Optical detection system for triazine based on molecularly-imprinted polymers. *Anal. Lett.* 30, 445-455.