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Transferrin-conjugated cationic liposome (Tf-liposome) was developed as a targeted gene delivery system by using heterobifunctional cross-linking agent, SPDP, and gradient metrizamide ultracentrifugation method. Physico-chemical properties of Tf-liposome were determined by scanning/transmission electron microscopy (SEM/TEM) and dynamic laser-light scattering method (DLS) with the mean diameter being  $584 \pm 15$  nm. Gel retardation assay was performed using various DDAB:DNA ratios and proved the 6:1 weight ratio formulation being the most compact with a slight positive zeta-potential. In vitro transfection was done in human cervical cancer cell line, HeLa, and the transfection efficiency of Tf-liposome was found to be 5-fold higher than that of un-conjugated (plain) DDAB liposome and 2-fold higher than that of Lipofectin™. Biocompatibility of Tf-liposome was also tested using human red blood cells (RBC) and their morphology remained unaffected after incubation with Tf-liposome at  $10 \mu\text{g/ml}$  concentration. In conclusion, a target-oriented gene delivery system of transferrin-conjugated cationic liposome (Tf-liposome) was made successfully and proved to be very efficient in DNA delivery into the cells in culture. Furthermore, its possible use as an *in vivo* gene delivery system is highly expected as suggested by its biocompatibility test using human RBC.

[PE1-12] [ 04/19/2001 (Thr) 15:30 - 16:30 / Hall 4 ]

#### The studies of interaction between methamphetamine and melanin pigment in hair

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There were many studies that suggested the amount and type of melanin present were major factors in determining how much drugs deposited in hair after exposure but the mechanism of drug entrapment in hair had been unknown. In vitro methamphetamine (MA)-melanin interaction was firstly studied using ultrafiltration membrane systems to interpret the deposition mechanism of methamphetamine in hair. The solutions of MA-melanin in amber vials were shaken by Recipro shaker and equilibrated at ambient temperature ( $20 \pm 0.5^\circ\text{C}$ ) for 24 hours. The concentrations of free drug were determined with HPLC systems. The binding parameters, association constant (K) and the number (n) of binding site per weight (mg) of melanin, were obtained from the Scatchard equation. The binding or association constant (K) and the number (n) of binding site of methamphetamine to melanin polymer were  $604 \text{ L/mole}$  and  $3.46 \times 10^{-5} \text{ M mg}^{-1}$ , respectively. This binding constant indicated that the interaction of methamphetamine to melanin was somewhat stronger than the published binding constants of some small molecules (p-toluene sulfonic acid (1), etc) to polymers (serum albumin, polyvinylpyrrolidone, etc). The Scatchard plot showed curvature at high concentrations of methamphetamine. This curvature usually indicated the existence of more than one type of binding site. The IR spectrum methamphetamine-melanin mixture showed band shift from  $3420 \text{ cm}^{-1}$  to  $3376 \text{ cm}^{-1}$  at N-H stretching region of methamphetamine. This shifting of the N-H stretching band of methamphetamine to lower frequency would be from hydrogen bonding with some groups (would be carboxyl or hydroxyl groups) of melanin. This MA-melanin interaction suggested that melanin as one component of hair was contributing to methamphetamine deposition in hair.

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#### Application of Vitamin E TPGS forming the solid dispersion with furosemide

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