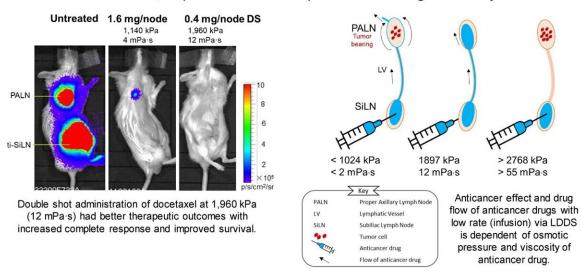


A novel approach for lymph node metastasis treatment

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Delivery of docetaxel via lymphatic network for Inhibition of Lymph Node Metastasis, Improves Tumor Response and Biological Safety



Osmotic pressure-and-viscosity-modified Docetaxel via lymphatic drug delivery system (LDDS) can inhibit tumor growth in the lymphatic system and prevent distant metastasis. Increases in osmotic pressure and viscosity lead to significant tumor inhibition in the sentinel lymph node (LN) and its' downstream LN. Moreover, we found the underlying mechanism of increased osmotic pressure and viscosity using Cisplatin (CDDP). The optimal range of osmotic pressure and viscosity solution covers all areas of its downstream LN via LDDS. Credit: Tohoku University



Researchers at Tohoku University have discovered a new approach for treating lymph node metastasis. Anticancer drugs are administered directly into the LNs under ultrasound guidance (lymphatic drug delivery system or LDDS) to target sentinel lymph nodes (LNs) and generate antitumor effects locally, preventing distant metastasis. This approach not only improves the anticancer effect but also reduces the nasty side effects commonly associated with systemic chemotherapy.

Details of the researchers' breakthrough are <u>published</u> in the journal *Biomedicine & Pharmacotherapy*. Professor Tetsuya Kodama from Tohoku University's Graduate School of Engineering led the research team.

Recent research has highlighted sentinel LNs as the gateway for metastasis. They are the initial lymph nodes that receive drainage from a primary tumor and indicate that metastasis has begun or could potentially occur. Thus, there is a need for improved control of LN metastasis without surgical or radiation interventions.

Still, the occurrence of side effects can differ from individual to individual and from one treatment to another. In fact, two individuals undergoing the same treatment may encounter markedly different experiences. Most commonly, some individuals experience uncomfortable side effects that gradually diminish over time.

In a prior investigation, some of the current researchers utilized LDDS incorporating heightened <u>osmotic pressure</u> and viscosity of docetaxel, yielding promising outcomes in the treatment of early-stage LN metastasis during preclinical research.

Based on this, they hypothesized that administering docetaxel—a widely used clinical anticancer drug—through LDDS at an osmotic pressure and viscosity of 1,960 kPa and 12 mPa•s would result in superior antitumor



effects, prolonged survival, and minimal adverse reactions.

To test this hypothesis, Kodama and his team administered the treatment to a preclinical model of a metastatic lymph node mouse model.

Their findings confirmed superior therapeutic outcomes, increased complete response, improved survival, and reduced adverse complications. The double-shot administration enhanced T helper cell differentiation in the spleen and sentinel LNs, emphasizing its potential in cancer management. Histopathological assessments confirmed the efficacy, highlighting the potential of LDDS in cancer management.

The research team emphasizes the safer nature of the new method. "LDDS is a groundbreaking method to improve treatment while decreasing side effects," says Ariunbuyan Sukhbaatar, assistant professor at the Graduate School of Dentistry and lead author of the paper. "This approach provides a safer alternative to LN removal and promises to improve cancer survival rates."

Meanwhile, Kodama says, "LDDS offers a novel, safe method for delivering <u>anticancer drugs</u> to susceptible LNs, presenting a promising approach for improving cancer treatment effectiveness while minimizing side effects."

Looking ahead, the research team aims to further investigate the LDDS with the goal of enhancing therapeutic responses against lymph.node metastasis. This will be achieved through clinical trials conducted in collaboration with medical professionals, aiming to validate the effectiveness of these approaches in cancer patients.

More information: Ariunbuyan Sukhbaatar et al, Docetaxel administered through a novel lymphatic drug delivery system (LDDS) improved treatment outcomes for lymph node metastasis, *Biomedicine &*



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