

This press release is an English translation of a Japanese-language press release. The official language of this press release is Japanese, and the Japanese version takes precedence over the English version in terms of content and interpretation.

<Press Release>

April 6th, 2024

**Publishment of data from our pipeline**  
**at the American Association for Cancer Research Annual Meeting 2024**

Chordia Therapeutics Inc. (“Chordia”), a biotech company engaged in the research and development of novel therapies for cancer, presented new data at the American Association for Cancer Research (AACR) Annual Meeting, April 5-10, 2024, in San Diego, California.

Regarding CTX-712, CLK inhibitor, a total of 46 patients with solid tumors and a total of 14 patients with hematologic malignancies were enrolled in the CTX-712-CL-01 study to evaluate the safety, pharmacokinetics (PK) / pharmacodynamics (PD) profile and preliminary efficacy as of data cutoff on November 20, 2023. Observed DLTs (Dose-limited Toxicity) were dehydration, platelet count decrease, and hypokalemia, and MTD (Maximum Tolerated Dose) was determined to be 140mg twice a week. Common any-grade-related adverse events were nausea and vomiting. Among all patients, partial responses were observed in 4 patients (4/46, 8.6%), who were all patients with ovarian cancers. (4/14, 28.6%). Among a total of AML and MDS patients (14), CR(Complete Remission) was observed in 3 patients, CRi (Complete Remission without incomplete hematologic recovery) was observed in 1 patient and MLFS (Morphologic Leukemia-Free State) was observed in 1 patient, so overall response rate is 42.9%. In PK analysis, a dose-dependent increase in systemic exposure of CTX-712 was observed, and a dose-dependent increase of exon skipping in two maker RNAs that are extracted from peripheral blood cell demonstrated PD effect induced by CTX-712.

Regarding CTX-439, CDK12 inhibitor, we reported (1) pharmacodynamics (PD) markers to monitor the suppression of CDK12/13 in humans, (2) patient selection biomarkers to determine sensitivity to CTX-439, and a significant anti-tumor effect was revealed in PDX models that showed sensitivity in the survey, (3) a synergistic effect in combination with PARP inhibitors and other chemotherapy and molecular-targeted drugs. in vitro and in vivo.

Regarding GCN2, we reported that our candidate compound inhibiting the HRI/PERK/GCN2 triple kinase in cells observed a distinct anti-growth effect for Multiple Myeloma with resistant to proteasome inhibitors when combined with proteasome inhibitors in vitro and in vivo.

## <Overview of presentation at AACR 2024>

### **Presentation on CTX-712, CLK inhibitor**

Abstract No	CT115
Title	A first-in-human phase I study of CTX-712 in patients with advanced, relapsed or refractory malignant tumors (CTX-712-CL-01 study): Efficacy and safety in a hematologic malignancies cohort

Abstract No	CT110
Title	A first-in-human phase I study of CTX-712 in patients with advanced, relapsed or refractory malignant tumors (CTX-712-CL-01 study): Efficacy and safety in solid tumor cohorts

Abstract No	2081
Title	Biomarkers for CLK inhibitor CTX-712 treatment response in myeloid neoplasms: Paving the way toward clinical trials

### **Presentation on CTX-439, CDK12 inhibitor**

Abstract No	3301
Title	Translational Research of CDK12/13 Inhibitor, CTX-439, Informing Clinical Trial Strategy

### **Presentation on GCN2 inhibitor**

Abstract No	1233
Title	Novel inhibitor targeting triple integrated stress response kinase HRI, PERK, and GCN2 provides new insights into overcoming resistance to proteasome inhibitors in multiple myeloma

### **About CTX-712**

CTX-712 is a first-in-class, orally available, and selective small molecule inhibitor of CDC2-like kinase (CLK), a key regulator of the RNA splicing process that plays an important role in cell growth. For more information, please visit: [jRCT2080224127](https://jRCT2080224127) for details on the Japan phase 1 clinical trial and [clinicaltrials.gov/](https://clinicaltrials.gov/) (NCT05732103) for details on the US Phase 1/2 clinical trial of CTX-712.

### **About Chordia Therapeutics**

Chordia is a clinical-stage biotech company based in Fujisawa, Kanagawa Prefecture, Japan, is developing novel cancer therapies. Chordia's lead asset, CLK inhibitor CTX-712, is currently in Phase 1 clinical study in Japan and Phase 1/2 clinical study in the US. CTX-712 potentially targets vulnerabilities in cancer and could benefit patients with various types of cancer. In addition to CTX-712, Chordia is researching several preclinical assets, including CTX-439, a CDK12 inhibitor, which might be effective in cancers with specific abnormalities, as well as GCN2 inhibitors.

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