

# uniQure

## Corporate Overview

Founded: 1998 in the Netherlands as Amsterdam Molecular Therapeutics (AMT)

Listed on NASDAQ: February 5, 2014 (QURE)

Full-time employees: Approximately 200 in Amsterdam, the Netherlands and Lexington, Massachusetts

Chief Executive Officer: Matthew Kapusta

**Our focus is to advance the future of medicine through gene therapy.**

**uniQure is delivering on the promise of gene therapy**, single treatments with potentially curative results. We have developed a modular technology platform to rapidly bring new disease-modifying therapies to patients with severe genetic diseases.

We have a focused pipeline of innovative gene therapies and have established clinical proof-of-concept in our lead indication, hemophilia B, and preclinical proof-of-concept in Huntington's disease. Our pipeline of adeno-associated virus (AAV)-based gene therapies has been developed using an innovative technology platform, supported by industry-leading proprietary commercial-grade manufacturing capabilities. Through recent collaborations and our strategic partnership with Bristol-Myers Squibb to develop gene therapies for cardiovascular diseases, we have taken the next steps toward developing gene therapies targeting chronic and degenerative diseases that affect larger populations.

**uniQure has built the world's leading gene therapy platform.**

**Our gene therapy technology consists of three key elements:**

1. A therapeutic gene cassette that carries a transgene that encodes for the expression of a therapeutic protein.
2. A best-in-class AAV-based vector delivery system for delivering the gene cassette. We use the AAV5 variant, or serotype, of vector in our product candidates in hemophilia B and Huntington's disease, and have exclusive, worldwide rights to AAV5 for use in therapeutic products delivered to the brain or liver.
3. Administration technologies to effectively deliver the relevant transgene into the liver or central nervous system.

Paired with our state-of-the-art manufacturing process, uniQure brings together all key elements for a successful gene therapy product.

## State-of-the-Art Manufacturing Capability

uniQure produces our AAV-based gene therapies in our own facilities using insect cells and baculoviruses, a common family of viruses found in invertebrates. uniQure has worked in insect cell baculovirus manufacturing for well over a decade and has established a very strong and unique patent position in fundamental aspects of the technology that are applicable to the entire industry.

Our facility in Lexington, Massachusetts is one of the largest, most versatile gene therapy manufacturing facilities in the world. uniQure has made significant investments in designing, constructing and equipping our 55,000 square-foot facility with state-of-the-art laboratories and commercial-scale production capabilities. The facility offers GMP production capabilities to support all of our existing programs, with flexibility to expand further.

## Three Key Areas of Focus, each with proof-of-concept

### Hemophilia B

uniQure is developing a gene therapy for hemophilia B, a severe orphan blood clotting disorder. Our gene therapy product candidate AMT-061 consists of an AAV5 vector carrying a gene cassette with the Padua variant of Factor IX (FIX-Padua). FIX-Padua expresses a protein that has been reported to provide an approximate eight to nine-fold increase in FIX activity compared to the wild-type FIX protein.

We believe AMT-061 has the potential to provide lower immunogenicity and durable, consistent and curative benefits to nearly all patients with severe and moderately severe hemophilia B. uniQure is currently enrolling patients in the Phase III HOPE-B pivotal study, and we expect topline data from our Phase IIb dose confirmation study by the end of 2018.

Clinical data from our Phase I/II trial demonstrate that our AAV5-based gene therapy has been safe, effective and durable, with no loss of efficacy at up to two years of observation. No patient in the study has had any loss of Factor IX (FIX) activity or capsid-specific, T-cell-mediated immune response.

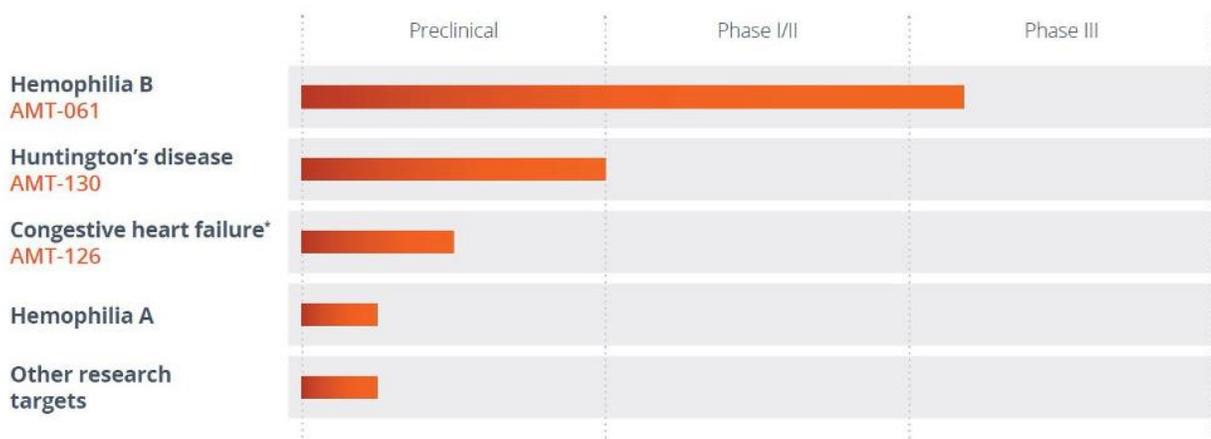
### Huntington's Disease

uniQure is developing a gene therapy for Huntington's disease, a rare, inherited neurodegenerative disorder that leads to loss of muscle coordination, behavioral abnormalities and cognitive decline, resulting in complete physical and mental deterioration over a 12- to 15-year period of time. Our product candidate AMT-130 consists of an AAV5 vector carrying an artificial micro-RNA specifically tailored to silence the huntingtin gene. The therapeutic goal is to inhibit the production of the mutant protein (mHTT) that causes the disease. Using AAV vectors to deliver micro-RNAs directly to the brain represents a highly innovative approach to treating Huntington's disease.

Data from multiple preclinical studies across five animal models show that a single administration of AMT-130 resulted in a dose-dependent and sustained reduction of mutant huntingtin protein in both the deep structures of the brain including the striatum and the putamen where Huntington's disease is known to manifest, and the cortex. We are very encouraged by the significant reductions in mutant huntingtin protein, and believe that knock-down of this magnitude has the potential to significantly alter the course of the disease. We expect to file an Investigational New Drug (IND) application in 2018 and to become the first AAV gene therapy to enter clinical development for Huntington's disease.

### Cardiovascular Disease

In April 2015, uniQure signed a strategic partnership with Bristol-Myers Squibb to discover and develop novel gene therapies to treat a range of cardiovascular diseases, initially focused on the one-time treatment of congestive heart failure (CHF) by selectively restoring cardiac deficiency of the calcium-binding protein S100A1, a master regulator of myocardial function. AAV-S100A1 has shown long-term therapeutic efficacy, safety and reduced mortality in a human-relevant, in vivo heart failure model compatible with clinical drug regimens. uniQure and Bristol-Myers Squibb continue to conduct IND-enabling studies for S100A1. In addition to S100A1, uniQure and Bristol-Myers Squibb have initiated work on three additional targets designated under the collaboration.



\*Partnered with Bristol-Myers Squibb