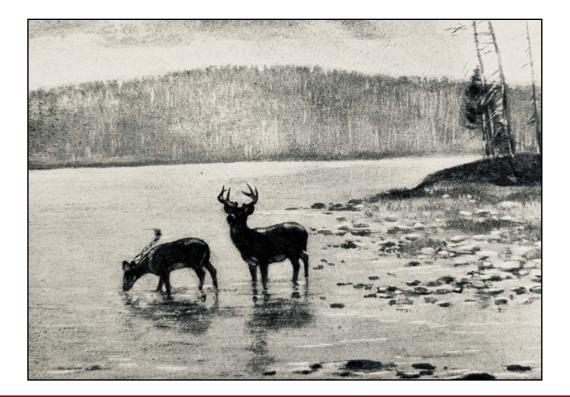
Chronic Wasting Disease: research and education update

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29 October 2019



CWD Research Activities





Research

- CWD requires an organized research effort
 - Bench-work and boots on the ground
- Organizing the Minnesota Center for Prion Research and Outreach (MNPRO)
 - Vision: multi-disciplinary center that strategically focuses on prion and proteinmisfolding diseases
 - Research hub for combating CWD and other neurodegenerative diseases (Alzheimer's, Parkinson's, ALS, etc.)

CWD: A pressing need for research Comparative medicine & biology Convening diverse expertise in animal health, cellular and protein biology, Chronic wasting disease ecological modeling, and human and animal protein-misfolding diseases (CWD) is a contagious, fatal neurological disease Think-tank environment affecting deer, moose, elk, Creating an incubator for cutting edge science and exploring new ideas to reindeer, and caribou. CWD is prion disease, Strategic research priorities and is similiar to Developing next-gen diagnostic tools, and leading the charge on bovine spongiform encephalopathy ("mad cow disease"), scrapie, and Coordination and collaboration Creutzfeldt-Jakob disease. Aligning efforts of multistate partners, and engaging stakeholders Currently, there is no evidence that CWD poses a risk for humans; however, the Centers for Disease Control and CWD has spread to 26 US states Prevention recommend since it was first detected in that people do not consume meat from Colorado in the 1960s animals known to be infected.

Minnesota Center for **Prion Research and Outreach**

College of Veterinary Medicine

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UNIVERSITY OF MINNESOTA Driven to Discover

Minnesota Center for Prion Research and Outreach

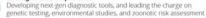
The Minnesota Center for Prion Research and Outreach (MNPRO) will be a multi-disciplinary center at the University of Minnesota focusing on the biology and epidemiology of human and animal prion diseases and related protein-misfolding disorders.

MNPRO will be a hub for combating neurodegenerative diseases, and will convene a range of U of M faculty and external collaborators to conduct research with a broad impact on protein-misfolding diseases such as Alzheimer's disease, Parkinson's disease, ALS, and emerging prion diseases such as chronic wasting disease.

A vision for research and outreach

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through research and outreach initiatives

Why the University of Minnesota

The University of Minnesota is uniquely prepared to address neurodegenerative diseases. As a comprehensive university with a diverse realm of expertise, the U of M has many of the necessary tools to safely and efficiently address these diseases. Our science is grounded by our location at the intersection of four environmental biomes, making our results relevant to a broad swath of North America.

Research

- MNPRO
 - Serve as a catalyst for fresh wave of prion research
 - Primary CWD research avenues
 - Next-generation diagnostic tools and biosurveillance
 - Environmental impact, transmission routes, and ecological modeling
 - Vaccines and therapeutics
 - CWD strain identification and zoonotic risk
 assessment
 - Establish CWD tissue biorepository





- Tiffany Wolf, DVM, PhD
 - Assistant Professor, Dept. Vet Population Medicine
 - Wildlife Epidemiologist
 - Working with Minnesota Grand Portage Band of the Lake Superior Chippewa Tribe to establish a Tribal CWD surveillance network
 - Recently awarded USFWS Tribal Wildlife Grant (~\$200k) to support this effort (2020 and beyond)





- Scott Wells, DVM PhD
 - Professor Dept. of Vet Population Medicine (epidemiologist)



- Three forthcoming papers on CWD exposures on 34 cervid farms (MN and WI).
- Major finding: 11 CWD-positive farms that were not considered high-risk farms (i.e., never imported CWD + animals) were located in areas near CWD + wildlife <u>link</u>



CWD Diagnostic Development Team



Dr. Schefers Dr. Larsen Dr. Skinner Dr. Seelig Dr. Oh

Collaborators: NIH Rocky Mtn Labs, Colorado State Univ Prion Research Center, Univ Of Maryland, Michigan State Univ, Public Health Agency of Canada, Alberta Centre for Prions and Protein Folding Diseases, Midwestern Univ Arizona



- CWD Diagnostic Development Team
 - Goal: develop advanced CWD diagnostics that are faster, more sensitive, and easier to use
 - Prototype(s) in 2 years
 - Functional with hunter harvested deer, live deer, and environmental samples
 - \$2M in July 2019 (\$259k Rapid Ag Response Fund, \$1.8M MN Legislature LCCMR fund)

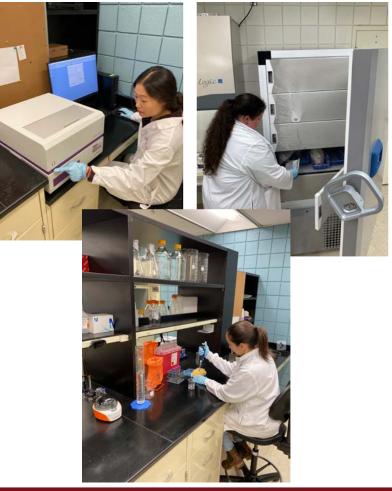


- CWD Diagnostic Development
 - Four research staff and three graduate students since July
 - Collaborating with DNR, BAH, Oxbow Park, hunters, etc. to secure tissues, fecal samples, etc.
 - Outfitted primary prion research laboratory
 - Cornerstone of MNPRO
 - Wet lab based in College of Veterinary Medicine



• MNPRO wet lab







- Dr. Skinner's Lab
 - Blood-based RNA biomarkers that can detect CWD infection



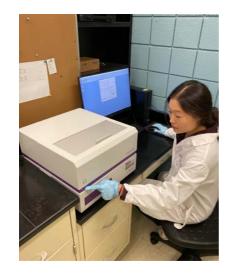
- Will test these RNA biomarkers in white-tailed deer of MN
- Primary Objective: Blood samples from recently harvested or live deer can be used to detect CWD infection





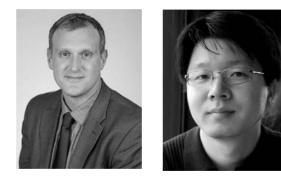
- Prion Research Lab: RT-QuIC functionality
 - Protein specialists trained at NIH Rocky Mtn. Labs and Colorado State Univ on RT-QuIC (Gage Rowden and Manci Li)
 - RT-QuIC: Diagnostic method capable of detecting CWD prions in tissues, blood, feces, soil, etc.
 - College of Vet Med lab will be first in the state of Minnesota to have RT-QuIC (November 2019)
 - Will use to screen over 500 deer sampled by the DNR (contains CWD + animals, blinded)
 - Diagnostic R&D focused on improving RT-QuIC







- CWD Diagnostic Development
- Antibody Engineering
 - New method of generating antibodies
 - Likely generate novel binding-affinities for CWD prions
 - Increased functionality and sensitivity for antibodybased diagnostics
 - Microfluidic experiments in Dr. Oh's laboratory ongoing



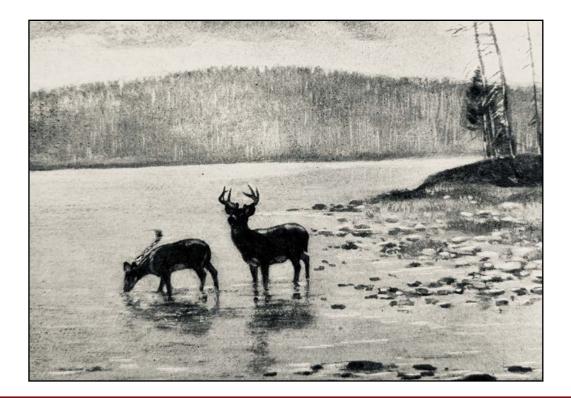




- Multiple groups across USA working on developing next-gen CWD diagnostics!
 - This is an area ripe for discovery, not a "moonshot"
- NIH Rocky Mtn Labs, USDA, USGS, Michigan State Univ, North Dakota State Univ, Univ of Texas, Univ of Minnesota, etc.

Michigan State University and Michigan Department of Natural Resources scientists are testing a faster, more accurate way to screen and diagnosis chronic wasting disease, or CWD, in deer. The three-year, \$900,000 project, funded by both institutions, will use RT-QuIC, a technology known to have better detection and sensitivity in real time.

Press release 30 Sept 2019





- College of Vet Med: Center for Animal Health and Food Safety (CAHFS)
 - CWD Watch website and educational materials
 - CWD Animation and videos
 - z.umn.edu/CWDWatch



Stay tuned for future conversations and interviews with Minnesota DNR scientists, the diagnostic development team, state legislators, Minnesota landowners, and human health experts.



What is CWD? >

Learn more about chronic wasting disease including which animals it affects, where it's found, and how it's spread.



CWD Progression >

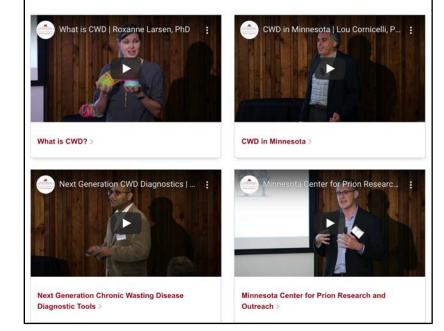
Chronic wasting disease can easily spread in nature, and it can take up to 2 years for CWD to kill an infected animal. But why is that?



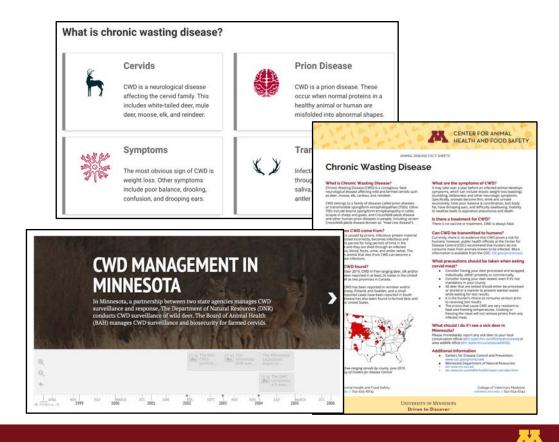
- CWD Watch website
 - UTalks (TED style talks from UMN experts)
 - Content will grow in coming weeks, months
 - z.umn.edu/CWDWatch

New! UTalks - Spotlight Science: Chronic Wasting Disease

U of M and Minnesota DNR scientists gathered on September 14th for the Spotlight Science event at the <u>Bell Museum</u>. Check out these UTalks from CWD researchers on key topics around the disease.



- CWD Watch website
 - Handouts, links to DNR, BAH, maps, etc.
 - Interactive CWD in MN timeline
 - z.umn.edu/CWDWatch



- Center for Infectious Disease Research and Policy
- CWD information, news, maps, etc.
- <u>http://www.cidrap.umn.edu/c</u>
 <u>wd</u>

CIDRAP Center for Infectious Disease Research and Policy

News & Perspective Infectious Disease Topics Antimicrobial Stewardship Ongoing Progra
 RENDING TOPICS Ebola Measles Antimicrobial Stewardship Chronic Wasting Disease

About CWD

Chronic Wasting Disease Resource Center

CWD Response, Research, and Policy Program

The Chronic Wasting Disease (CWD) Response, Research, and Policy Program addresses the transmission of CWD in cervids and its potential for spread to humans and other animal species. The program supports current and reliable information on CWD for the public, including hunters; the medical, veterinary and public health communities; wildlife scientists and managers; and public policymakers.

About CIDRAP's CWD Program



Expert Advisory Group

The program includes 49

About CWD

Chronic wasting disease (CWD) is a prion disease that affects several cervid species: doer, elk, reindeer, sika doer, and moose. CWD was first identified in 1967 in a captive mule deer living in a Colorado research fallity. In 1981, CMD was detected for the first time in a wild cervid. Since these initial detections, CWD has been identified in 36 states and three Canadian provinces. It has also been detected in Finland, Norway, South Knore, and Sweden.

CWD is believed to be transmitted horizontally (i.e., animal-to-animal contact) through indictious body fluids such as alloy, urine, and foces. Once accurated into the environment, CWD prions can persist for years and withstand extremely high levels of disinfectants such as heat, radiation, and formaldelyde. CVD prions also are capable of binding to certain plants, with the ability to be transported while still remaining infectious. CVD is increasing in cervida as more animals come into contact with infections prions, usually via direct contact with an infected cervid and its bodyl fluids, although viable CVDP prions in the environment can also infect animals. As more cervida become infected, the frequency of these exposures and subsequent environmental contamination grows. Evidence also suggests that vertical transmission (i.e., parent to offspring) can occur, although is overall impact on the ecology of CVDP is not entirely understood at this inte.

Since CWD is now an established widlife disease in North America, proactive steps, where possible should be taken to limit transmission of CWD among animals and reduce the potential for human exposure. Although CWD has not yet been found to cause infections in humans, numerous hashi genricis have taken the stance tata people should not be consuming CWD-positive animals. Since 1997, the World Health Organization has recommended that agents of any priori disease should more enter the human food chain. Likewise, the US Centers O Foisses Control and Prevention, Health Canada, and multiple provincial and state health and natural resources agencies recommend that people should not consume the mean of an aming found to be positive for CWD.

Given the typical ten year or longer incubation period of prion-associated conditions, improving public health measures now to proven thuman exposures to CVD prions and to further understand the potential risk to humans may reduce the likelihood of an event like borine spongiform encephalopathy (BSI). In BSE, also known as "mad coor diseases, some Brittish officials in the 1990s declared there was no risk of transmitting BSE prions through the consumption of contaminated bee, only to confirm related human aces of a similar prion disease in the neuring years.

• Augmented Reality CWD Displays (College of Vet Med and Center for Animal Health and Food Safety)



Bell Museum (14 Sept 2019)





- Augmented Reality CWD Displays
 - Portable, easy to setup, travel around the state!



Winona State Univ. (10 Oct 2019)

Phone / Tablet App to interact with display for more info









- Bell Museum of Natural History CWD Event (14 Sept)
 - Organized by CAHFS
 - DNR, BAH, Elk Breeders Assoc.
 - TED style talks, interactive displays, facilitate CWD discussion
 - Talks available at:
 - z.umn.edu/CWDWatch



- CWD Seminars
- Prion biology and the science of CWD
 - CVM Agents of Disease
 - Eagle Bluff Nature Center
 - Red Wing Rotary Club
 - Winona State University
 - 4 planned in coming months

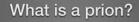
Chronic Wasting Disease in SE Minnesota

September 16, 2019 by Rich Wicks – Leave a Comment



Summary: Education and Outreach

- Critical need for additional prion science
 outreach!!!
- Misunderstanding and fear of CWD
 - Causative agent
 - Not a bacteria or a virus, but a prion
 - What is a prion?
 - Risk to humans
 - Requires understanding of diversity of prion diseases
 - No known human transmission but there is risk! Why?



- All mammals have prion proteins
- Normal cellular function
 - Copper and metal processing



Important functional roles
 in nerve cells







Potential confusion with respect to CWD...

Emerging virus impacting deer: Epizootic Hemorrhagic Disease (EHD)



EHD and CWD: What's the Difference?

Hemorrhagic Disease (EHD, Bluetongue)

Basics EHD: Viruses spread by biting gnats in late summer. Symptoms include fever and internal hemorrhaging.

Pathway EHD: Cannot be spread from deer to deer, only through bites from infected insects.

Victims EHD: Bucks and does of all ages are equally susceptible to being bitten by infected insects.

Location EHD: The viruses are present everywhere in North America, but outbreaks are associated with drought and extreme heat, usually in late summer.

Chronic Wasting Disease (CWD)

CWD: A syndrome of the central nervous system in which the brain deteriorates. Caused when normal proteins called "prions" become deformed.

CWD: Spread deer-to-deer through direct contact, or contact with the saliva, urine, feces, blood, and body parts of infected deer or infectious materials in soil.

CWD: Higher infection rates among bucks, particularly mature bucks, most likely because they cover more ground and contact other deer more often.

CWD: Present in deer or elk herds in 23 states and two Canadian provinces. Preventing CWD's spread to new areas is critical. Transportation of live, infected deer/elk or their parts is the primary long-distance pathway.

Credit: Quality Deer Management Association



EHD vs CWD

Mortality rate	EHD : Some deer survive infection. Herd immunity/survival is higher in areas with longer historical exposure.	CWD: Always fatal.
Speed of death	EHD : For those deer that die, death usually occurs within a few days of infection.	CWD : Incubates in infected deer for an average of around one to two years before symptoms appear. During incubation, deer can spread CWD to other deer.
Durability	EHD : Viruses cannot survive outside the bodies of the insect vector or the deer/ elk host.	CWD : Infectious materials remain viable indefinitely (years) in the envionment and are shed in feces, urine, saliva, blood and carcasses of infected animals.
Human health	EHD: Cannot infect people, either through insect bites or by consuming infected deer.	CWD : No evidence that it is a health issue in humans, but the Centers for Disease Control and Prevention urges caution in handling venison in infected areas, and suggests hunter-harvested deer be tested for CWD before being consumed.
Long-term	EHD: Outbreaks vary locally from mild to serious, but deer populations rebound. Whitetails have lived with and adapted to these viruses for decades.	CWD : Infection rates at some outbreak sites are climbing slowly but steadily, and the long-term impact is still not clear. Over time, CWD may alter the social structure of deer herds by eliminating mature animals.

Credit: Quality Deer Management Association



Recent CWD Publications

- Reviews
 - <u>CWD: current assessment of transmissibility (Sakudo, Cur Issues</u> <u>Mol Biol)</u>
 - <u>CWD: emerging prions and their potential risk (Hannaoui et al,</u> <u>PLOS Pathogens)</u>
 - <u>CWD in Cervids: prevalence, impact and management strategies</u> (<u>Rivera et al., Vet Med</u>)
- Opinion/Hypothesis
 - <u>CWD in Cervids: implications for prion transmission to humans and</u> other species (Osterholm et al., MBIO)



Recent CWD Publications

- Research
 - Inactivation of CWD prions using sodium hypochlorite (Williams et al., PLOS One)
 - <u>Detection of CWD in cervids by RT-QuIC assay of third eyelids</u> (Cooper et al., PLOS One)
 - <u>Sodium hydroxide treatment effectively inhibits PrPCWD</u> replication in farm soil (Sohn et al, Prion)
 - <u>Insights into the bidirectional properties of the sheep-deer prion</u> transmission barrier (Harrathi et al., Mol Neurobiol)



Recent CWD Publications

- Research cont.
 - <u>Rapid recontamination of a farm building occurs after attempted</u> prion removal (Gouch et al, Vet Rec)
 - <u>Transmission studies of CWD to transgenic mice overexpressing</u> <u>human PrP using the RT-QuIC assay (Race et al., Vet Res)</u>
 - <u>Spatial heterogeneity of prion gene polymorphisms in an area</u> recently infected by CWD (Miller and Walter, Prion)



Thank you!

- Minnesota Agricultural Experiment Station: Rapid Agricultural Response Fund
- Minnesota Legislature, LCCMR
- Grand Portage Band of Chippewa, USFWS
 - Seth Moore and Tiffany Wolf



- AGREETT (Agricultural Research, Education, Extension and Technology Transfer Program)
- DNR, BAH, Elk Breeders Association, Oxbow Park and Zollman Zoo





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