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#### Covid-19 Clinical Update 4/9/2020

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# COVID-19 UPDATE

HANA AKSELROD, MD, MPH GW DIVISION OF INFECTIOUS DISEASES 4/9/2020 1. EPIDEMIOLOGY

2. PATHOPHYSIOLOGY

3. TESTING

4. TREATMENT

5. GW UPDATES

https://www.smithsonianmag.com/smart-news/tiger-bronx-zoo-tested-positive-covid-19-180974624/

# EPIDEMIOLOGY

### Known coronavirus cases in D.C., Maryland and Virginia

There are a total of **10,634** reported cases in the region.



#### Total cases reported in the District, Maryland and Virginia

New cases each day

https://www.washingtonpost.com/graphics/local/dc-maryland-virginia-coronavirus-cases/ https://www.washingtonian.com/2020/04/03/dcs-covid-19-cases-are-younger-and-more-female-than-expected/



# PATHOPHYSIOLOGY



## **ACE-2 Receptors**

- Does ACE2 expression underpin COVID-19 severity?
- Angiotensin-converting enzyme 2 (ACE2)
  - Co-opted by SARS-CoV-2 to enter epithelial cells
  - Expression declines with age
  - Upregulated in patients treated with ACEi/ARB
- Anti-inflammatory effects
  - angiotensin 2  $\rightarrow$  angiotensin 1-7
  - angiotensin 1  $\rightarrow$  angiotensin 1-9
- Would ACEi treatment help compensate for age-related loss of ACE2 receptor and reduce severity of illness?
  - Lower mortality observed from other ILI/PNA/ARDS?
  - Any benefit to initiating ACEi?

#### April 3, 2020

## The Dilemma of Coronavirus Disease 2019, Aging, and Cardiovascular Disease

#### Insights From Cardiovascular Aging Science

Majd AlGhatrif, MD, MA<sup>1,2</sup>; Oscar Cingolani, MD<sup>3</sup>; Edward G. Lakatta, MD<sup>1</sup>

 $\gg$  Author Affiliations | Article Information

JAMA Cardiol. Published online April 3, 2020. doi:10.1001/jamacardio.2020.1329

#### April 3, 2020

#### Coronavirus Disease 2019 (COVID-19) Infection and Renin Angiotensin System Blockers

Chirag Bavishi, MD, MPH<sup>1</sup>; Thomas M. Maddox, MD, MSc<sup>2,3</sup>; Franz H. Messerli, MD<sup>4,5,6</sup>

#### $\gg$ Author Affiliations ~~ | Article Information

JAMA Cardiol. Published online April 3, 2020. doi:10.1001/jamacardio.2020.1282



### From: The Dilemma of Coronavirus Disease 2019, Aging, and Cardiovascular Disease: Insights From Cardiovascular Aging Science

JAMA Cardiol. Published online April 03, 2020. doi:10.1001/jamacardio.2020.1329



#### Figure Legend:

Schematic of Inflammatory Profile Before and After Coronavirus Disease 2019 (COVID-19) InfectionSimplified schematic of the preinfection inflammatory profile among predisposed older individuals vs their younger counterparts. ACE2 indicates angiotensin-converting enzyme 2; CVD, cardiovascular disease; SARS-CoV-2, severe acute respiratory syndrome coronavirus 2.

#### DATE OF DOWNLOAD: 4/8/2020

## Can My Pet Get COVID-19?

- SARS-CoV-2 replicates poorly in dogs, pigs, chickens, and ducks, but effectively in **ferrets** and **cats**
- Symptoms: loss of appetite, fever
- Inoculated intranasally & exposed in adjacent cages
- Viral RNA recovered from tonsils, palate, trachea, lung tissue, and feces
- 3/3 inoculated cats and 1/3 exposed cats developed detectable viral RNA and later antibodies against SARS-CoV2
- "It was difficult to perform regular nasal wash collection on the subadult cats because they were aggressive."

#### REPORT

### Susceptibility of ferrets, cats, dogs, and other domesticated animals to SARS-coronavirus 2

Jianzhong Shi<sup>1,\*</sup>, Zhiyuan Wen<sup>1,\*</sup>, Gongxun Zhong<sup>1,\*</sup>, Huanliang Yang<sup>1,\*</sup>, Chong Wang<sup>1,\*</sup>, Baoying Huang<sup>2,\*</sup>, Renqiang Liu<sup>1</sup>, Xi... + See all authors and affiliations

Science 08 Apr 2020: eabb7015 DOI: 10.1126/science.abb7015



# TESTING

## **Clinical Performance of Tests**

Figure. Severe Acute Respiratory Syndrome Coronavirus 2 Distribution and Shedding Patterns Among 20 Hospitalized Patients



## Testing

- Rapid antibody binding tests
- "Certifying immunity"
- Wide-scale testing in UK
- Problems with false-negative and false-positive results
  - Validated against ELISA against S protein and other viral proteins
  - Compared against prior donated samples of blood by same individuals
- Cross-reactivity with antibodies against seasonal coronaviruses
- Projected timeline to reliable test: "At least a month"



DIAGNOSTIC TOOLS

CORONAVIRUS

SHARE TH



**D** rofessor Sir John Bell explores the challenges of antibody testing for Covid-19, noting that no tests to date have performed well.

# TREATMENT



#### Available online 20 March 2020, 105949 In Press, Journal Pre-proof (?)

International Journal of Antimicrobial Agents



Hydroxychloroq treatment of CO

HEALTH

### label non-rando

Philippe Gautret <sup>a, b, \$</sup>, Jean-Christop Meddeb<sup>a</sup>, Morgane Mailhe<sup>a</sup>, Barbari Vieira<sup>a</sup>, Hervé Tissot Dupont<sup>a, c</sup>, Stér Scola <sup>a, c</sup>. Jean-Marc Rolain <sup>a, c</sup>. Philip 

### FRENCH HOSPITAL STOPS **HYDROXYCHLOROQUINE TREATMENT FOR COVID-19 PATIENT OVER MAJOR CARDIAC RISK**

**Retraction Watch** Tracking retractions as a window into the scientific process

Intimicrobial

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standard"

ne and

Hydroxychloroquine-COVID-19

-label non-



does not meet the [International Society of Antimicrobial *Chemotherapy's] expected standard, especially relating to the* lack of better explanations of the inclusion criteria and the triage of patients to ensure patient safety.

### AHA, ACC, HRS Caution Use of COVID-19 Therapies Hydroxychloroquine and Azithromycin in Cardiac Patients

*New COVID-19 drug therapy guidance from the American Heart Association, the American College of Cardiology and the Heart Rhythm Society* 

- Withhold the drugs in patients with baseline QT prolongation (QTc ≥500 msec) or known congenital long QT
- Monitor cardiac rhythm and QT interval; withdraw the drugs if QTc exceeds 500 msec
- In patients critically ill with COVID-19 infection, frequent caregiver contact may need to be minimized, so
  optimal electrocardiographic interval and rhythm monitoring may not be possible
  - ECG preferred
  - Continuous telemetry monitoring as alternative
- Correct hypokalemia to levels of >4 mEq/L
- Correct hypomagnesemia to levels of >2 mg/dL
- Avoid other QTc prolonging agents whenever feasible

**Table.** Torsade de pointes potential and post-marketing adverse events associated with possible

 COVID-19 repurposed pharmacotherapies.

Possible COVID-19 Treatment	CredibleMeds	VT/VF/TdP/LQTS	Cardiac Arrest in
	Classification	in FAERS	FAERS
Repurposed antimalarial agents			
Chloroquine	Known risk	72	54
Hydroxychloroquine	Known risk	222	105
Repurposed antiviral agents			
Lopinavir/ritonavir	Possible risk	27	48
Adjunct agents			
Azithromycin	Known risk	396	251
OVID-19 indicates coronavirus disease 2019: FAFRS, US Food and Drug Administration Adverse			

COVID-19 indicates coronavirus disease 2019; FAERS, US Food and Drug Administration Adverse Event Reporting System; LQTS, long QT syndrome; and TdP, torsade de pointes.





# GW UPDATES