

**What will you die of?
&
Health in 2030**

Week 4 Lecture 2

Before we go down the technology rabbit hole...

We need to talk about the economics of technology

Otherwise, very little makes sense

Newspaper headlines

Starvation

Global warming

High drug prices

New dangerous diseases

And more...

Global warming (bio)tech

If you are a (bio)engineer, then in three minutes, you

can think of ways to address global warming:

- Sunlight and CO₂ to biofuels
- Use Photovoltaics to split water, make Hydrogen
- Collect CO₂ from atmosphere and store deep underground

water, and a genetically modified bacterium.

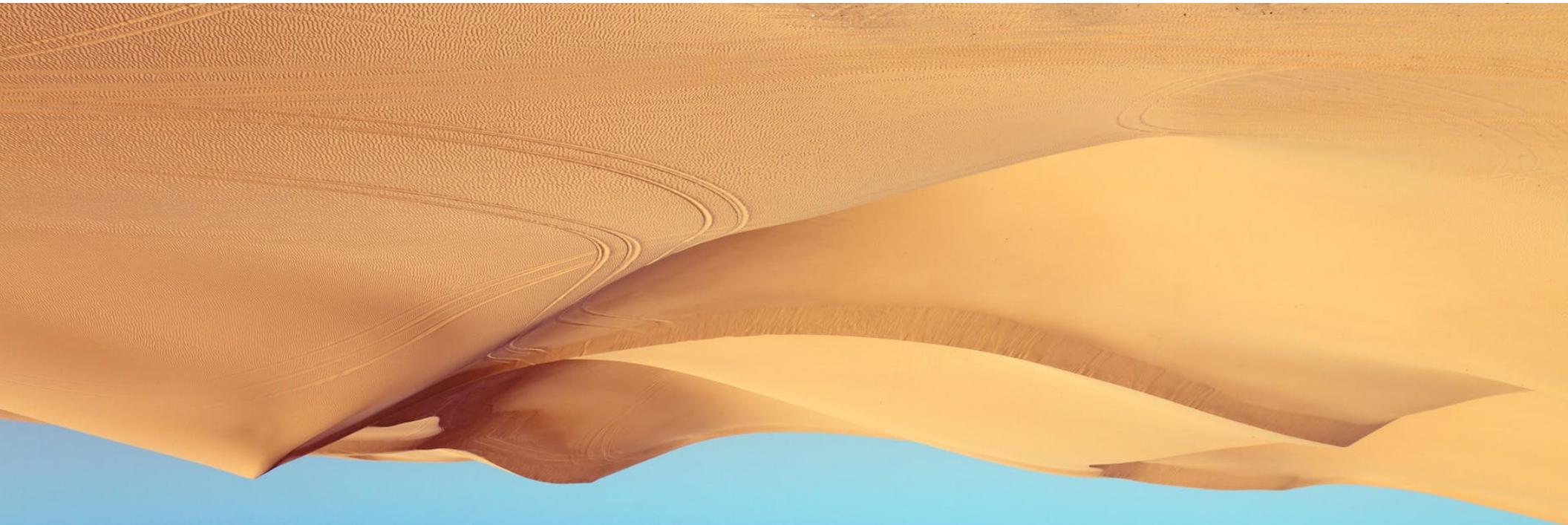
Joule was designing a system that would produce diesel fuel or gasoline using nothing more than the sun, carbon dioxide,

COURTESY OF JOULE UNLIMITED



How a biofuel dream turned into a nightmare

SCOTT KIRNSNER | INNOVATION ECONOMY



There is one tiny little problem

Deals

Aramco Pumps Oil at Fraction of Rivals' Costs and May More of It

By Anthony Dipaola

April 1, 2019, 6:24 AM PDT

Updated on April 1, 2019, 2:00 PM PDT

- ◀ Cost of extracting crude is \$2.80 a barrel; bond prospects
 - ◀ Production is more than 5 international oil companies com

\$2.80 a barrel:

Note: Some data provided by person who saw presentation for investors

Global warming tech



1 barrel = 42 gallons

42 gallons at the pump

= \$189

$\$189 / 2.80 =$
6650% profit



Global warming tech

56% profit



100 fold difference



6650% profit

Cost to extract crude from
the ground
\$0.06 per gallon



Cost to make 'crude' at the plant
\$3 per gallon

https://www.card.iastate.edu/research/biorenewables/tools/hist_bio_gm.aspx



Global warming tech

What's the
issue with this
calculation?

Compared to biofuels, the refining of petroleum is less expensive as it is highly optimised and nothing is wasted. Projections of price and technology development that advanced biofuels could be competitive with fossil fuels on a volume basis, but timing ranges from 2020-2030 and at an oil price ranging from US\$70-US\$150/barrel. At a low price of US\$60/barrel only conventional biofuels such as sugar cane ethanol can currently compete directly on a volume basis.
As technology advances, biofuels can come to the market at lower costs, with a lower environmental impact and higher GHG reductions. As seen with conventional biofuels, a cost reduction of between 1.5-3 times can be achieved in 10-20 years.

<http://www.biofuelsforurope.eu/cost-competitiveness/>

Prediction - Biofuels will compete with
Aramco in 90 years assuming production cost
goes down by 1/2 every 20 years

2030 - \$50
2050 - \$25
2070 - \$12.5
2090 - \$6.25
2110 - \$3.13



Global warming tech

Competitiveness: Biofuels vs Petroleum-based fuels

Newspaper headlines

Starvation

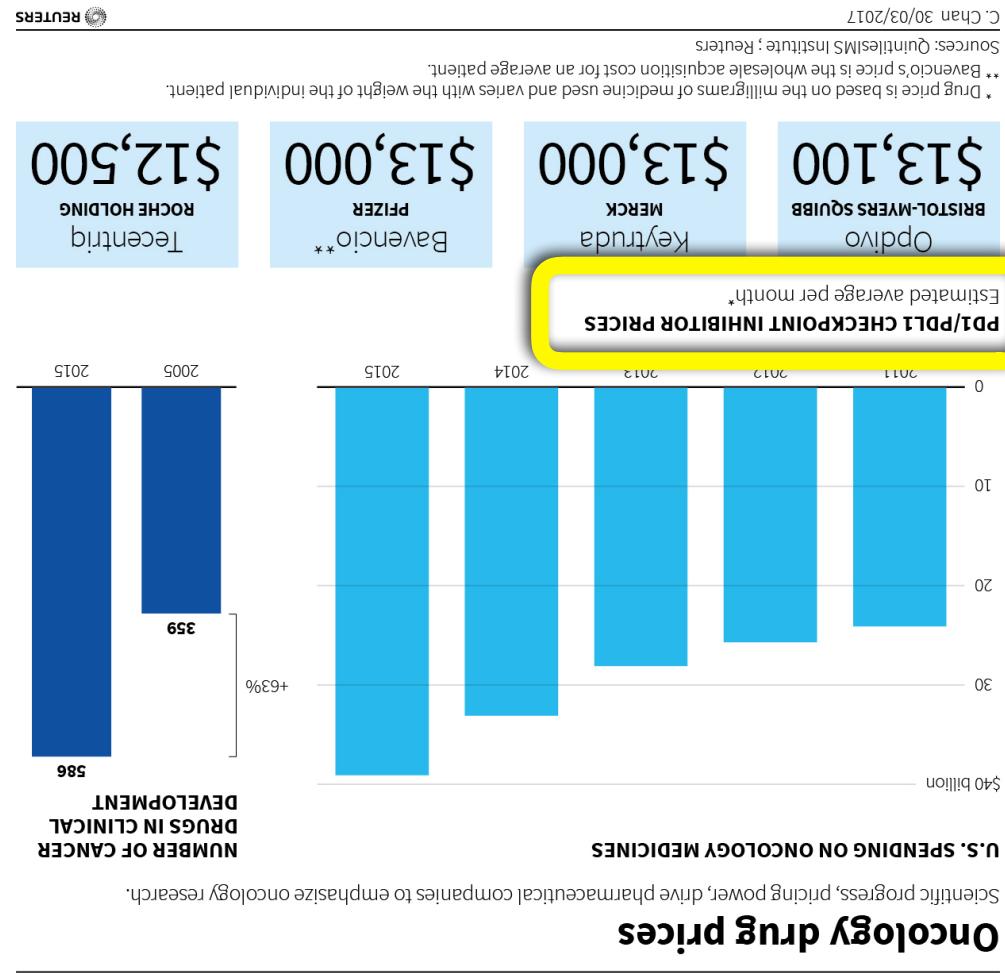
Global warming

High drug prices

New dangerous diseases

And much more...

Drug (bio)tech



The Genius Issue

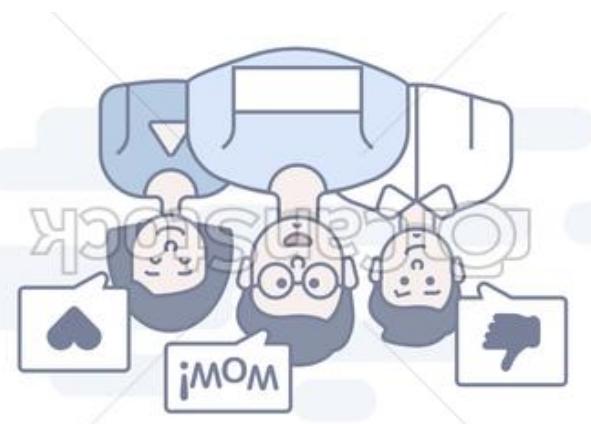


TIME

Jan. 2013

Double Issue

everyone can afford!
make cheap drugs that
use biotechnology to



Drug (bio)tech

Drug (bio)tech

Let's look at manufacturing costs...

Words that do not appear in the report:

Cost of goods sold (GOCs)

Materials

Manufacturing cost

Appendix A
2018 Financial Report

Drug (bio)tech

(Labor, packaging, shipping,
quality control, chemicals)
Cost of sales: 21%

Balipark:
Cost of drug
chemicals, relatively
small part of COs

Marketing: 27%
R&D: 14%

(MILLIONS OF DOLLARS)	2018	2017	2016	18/17	17/16	Year Ended December 31,	% Change
Revenues	\$ 53,647	\$ 52,546	\$ 52,824	2	—		
% of revenues	11,248	11,228	12,322	(1)	(9)		
Selling, informational and administrative expenses	14,455	14,804	14,844	(2)	—		
% of revenues	26.9%	28.2%	28.1%				
Research and development expenses	8,006	7,683	7,892	4	(3)		
% of revenues	14.9%	14.6%	14.9%				
Amortization of intangible assets	4,983	4,758	4,056	3	17		
% of revenues	9.1%	9.1%	7.7%				
Restocking charges and certain acquisition-related costs	1,044	351	1,565	*	(78)		
% of revenues	1.9%	0.7%	3.0%				
Other (income)/deductions—net	2,116	1,416	3,794	49	(63)		
% of revenues	4.1%	3.0%	1.0%				
Income from continuing operations	11,885	12,305	8,351	(3)	47		
Effective tax rate	22.2%	23.4%	15.8%				
Provision/(benefit) for taxes on income	706	(9,049)	1,123	*	*		
Effective tax rate	5.9%	(73.5)%	13.4%				
Income from continuing operations	11,179	21,353	7,229	(48)	*		
Discounted operations—net of tax	20.8%	21.355	7,246	*	(87)		
Net income before allocable to noncontrolling interests	11,188	21,355	7,246	(48)	*		
Less: Net income attributable to Pfizer Inc.	36	47	31	(24)	54	*	
% of revenues	20.9%	40.6%	13.7%				
Net income attributable to noncontrolling interests	36	47	31	(24)	54	*	
% of revenues	20.9%	40.6%	13.7%				
Net income attributable to Pfizer Inc.	\$ 11,153	\$ 21,308	\$ 7,215	(48)			

ANALYSIS OF THE CONSOLIDATED STATEMENTS OF INCOME

quality control, chemicals)

(Labor, packaging, shipping,

Cost of sales: 21%

Marketing: 27%
R&D: 14%

So - before you invest in, or start, a business
in bio/tech/health, run the numbers - what
are the basics of the business, and what does
the completion look like?

Today's goal: Examine 2 case studies at the cross section of biengineering and health.

Social

Key: Opportunities can fail for many reasons, **Technical** and

just because you want something to come true, doesn't make it come true.

How can we imagine and realize the future?



Case#1 Artemisinin: Low cost & Local production of high value chemicals

Malaria is one of the most severe public health problems worldwide. A leading cause of death and disease in many developing countries, where young children and pregnant women are the groups most affected.

According to the [World Health Organization's World Malaria Report 2013](#) and the Global Malaria Action Plan

- 3.2 billion people (half the world's population) live in areas at risk of malaria transmission in 106 countries and territories

- In 2012, malaria caused an estimated 207 million clinical episodes, and 627,000 deaths. An estimated 91% of deaths in 2010 were in the African Region.

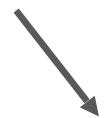
Centers for Disease Control and Prevention

CDC 24/7: Saving Lives, Protecting People™



Case-I: Malaria

Policy, economics



People could actually access it?

and

There were a good antimalarial drug,



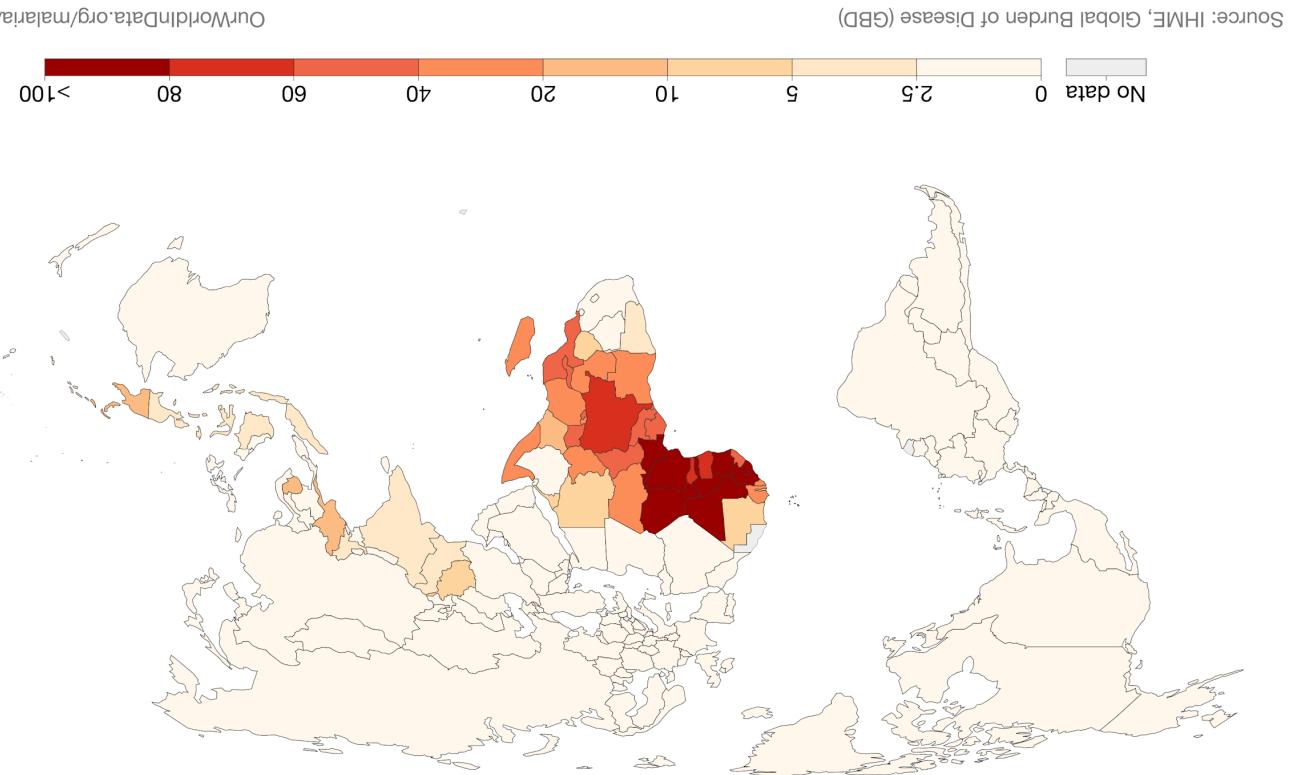
It would be nice if....

Science

Case background - Malaria death rates (per 100,000)

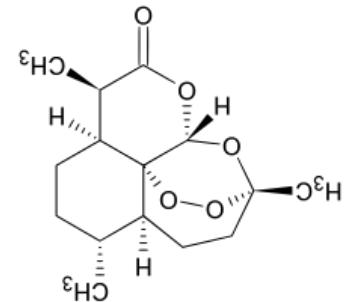
Malaria death rates (per 100,000), 2016

Age-standardized death rates from malaria, measured as the number of deaths per 100,000 individuals.
Age-standardization assumes a constant population age & structure to allow for comparisons between countries
and with time without the effects of a changing age distribution within a population (e.g. aging).



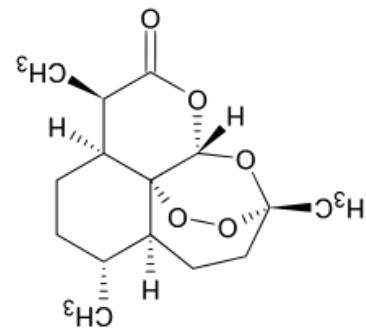
Case background - Artemisinin

- Artemisinin is isolated from the plant *Artemisia annua* (sweet wormwood) - (Tu Youyou received The Nobel Prize in Physiology or Medicine 2015)
 - Used to treat malaria and parasitic worm infections
 - Advantage: kills parasites faster and all the stages of life cycle
 - Disadvantage: low bioavailability, and high cost
 - Concern: Use of the drug by itself is explicitly discouraged by the WHO. Signs of resistance by malarial parasites



amyris

Case background - Artemisinin



Alternative approach:

No farming
Double the global supply
Stabilize price to \$300/kg.
Prices will drop quickly - tens of
dollars per kg.
Can be produced anywhere in
the world, rapidly

Almost zero land use
No solvents waste

Grow wormwood, thousands of
farmers and 20,000 hectares
in Kenya, Tanzania,
Madagascar, Mozambique, India,
Vietnam, and China
Extract compound via solvents (diethyl ether)
\$100-1000 per kg (limited supply)

Farming approach:





Production of the antimalarial drug precursor artemisinic acid in engineered yeast

Letter | Published: 13 April 2006

(source)

Nature 440, 940–943 (2006) | Download Citation ↗
Withers, Yochirō Shiba, Richmand Sharpone & Jay D. Keasling ↗
Kimbrell A., Ho, Rachel A., Eadhus, Timothy S., Ham, James Kirby, Michelle C. Y. Chang, Snyder T.,
Dae-Kyun Ro, Eric M. Paradise, Mario Ouellet, Kart J. Fisher, Karyn L. Newman, John M. Nduagu,
Desai, R. & Keasling, J. D. (2006). Production of artemisinic acid in yeast. *Nature*, *440*, 940–943.

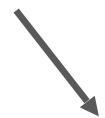
“Here we report the engineering of *Saccharomyces cerevisiae* to produce high titres (up to 100 mg l⁻¹) of artemisinic acid to synthesize an engineered mevalonate pathway ... The synthesized artemisinic acid is transported out and retained on the outside of the engineered yeast, meaning that a simple and inexpensive purification process can be used to obtain the desired product.

Although the engineered yeast is already capable of producing artemisinic acid at a significantly higher specific productivity than *A. annua*, yield optimization and industrial scale-up will be required to raise artemisinic acid production to a level high enough to reduce artemisinin combination therapies to significantly below their current prices.”

What is the science behind the approach?

Case-I: Malaria

Policy, economics



People could actually access it?

and

There were a good antimalarial drug,



It would be nice if....

Science

“In the constant fight between microbes and people, attempts to rein in the malaria parasite have just taken an interesting turn. On Thursday the founder of Amryris Biotech triumphantly announced production of 70m doses of the anti-malarial compound artemisinin. This sounds like good news for poor people but may be a step backwards — the start of a new high-tech assault on farmers.”

Source

▼ A farmer harvests sweet wormwood trees in Youyang, rural Chongqing, China, 2006. Photograph: Michael Reynolds/EPA



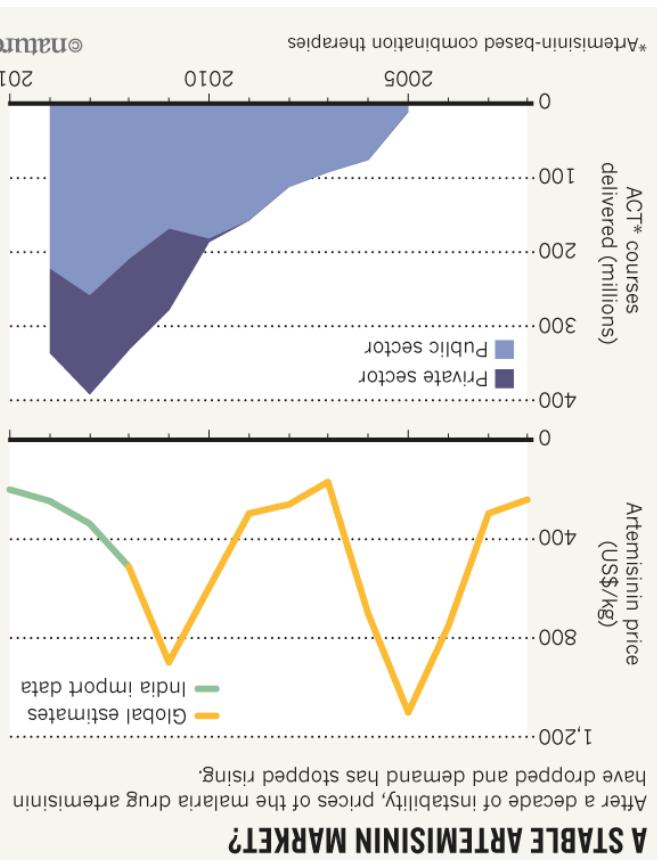
Artemisinin breakthrough by synthetic biologists threatens to open new front in battle between microbes and people

Synthetic anti-malarial compound is bad news for artemisia farmers

What are the **societal** aspects of approach?

Synthetic biology's first malaria drug meets market resistance (2016)

Commercial use of genetically engineered yeast to make medicine has modest impact.



"That is partly because of a glut in agricultural artemisinin. For the past two years, the naturally derived chemical has sold for less than \$250 per kg ... "If that price is already very low and there's a bumper crop, there's no reason to fire up a fermenter," says Jay Keasling of the UC Berkeley, who led the team that first developed the yeast strain... "I'd like to see semi-synthetic artemisinin take over as the dominant form, and some day I think it will," says Keasling. "But we have to be patient,"

Source

Share back to the entire class

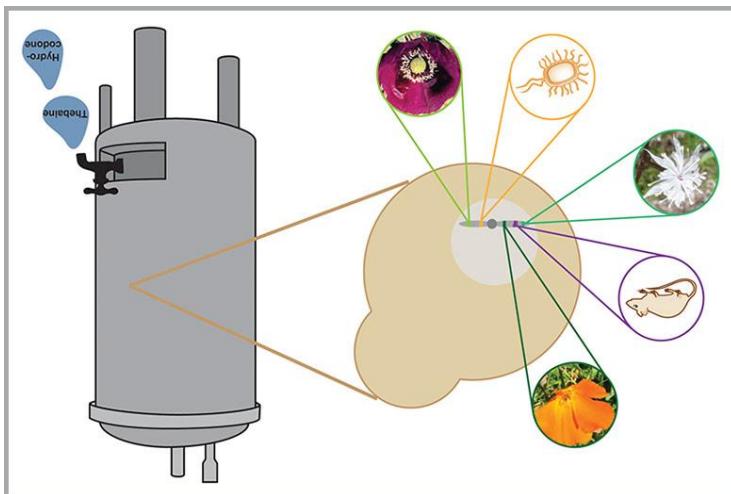
1. Who are the key stakeholders in this case?
2. Was Amyris successful? Why? What (if anything) went wrong?
3. If you were in charge of Amyris, what would you have done differently?
4. Are there any concerns associated with easy (affordable) access to Artemisinin?

In small groups discuss the following:

Activity-1: Discussion in Groups (2 mins)

Case#2 Opioids: Low cost & Local production of high value chemicals

(Image credit: Stephanie Galanis, Smolke Lab)



Stanford researchers genetically engineer yeast to produce opioids from plants, but Christina Smolke and colleagues have genetically modified yeast to make it in just a few days. The technique could improve access to medicines in impoverished nations, and later be used to develop treatments for other diseases.

AUGUST 13, 2015



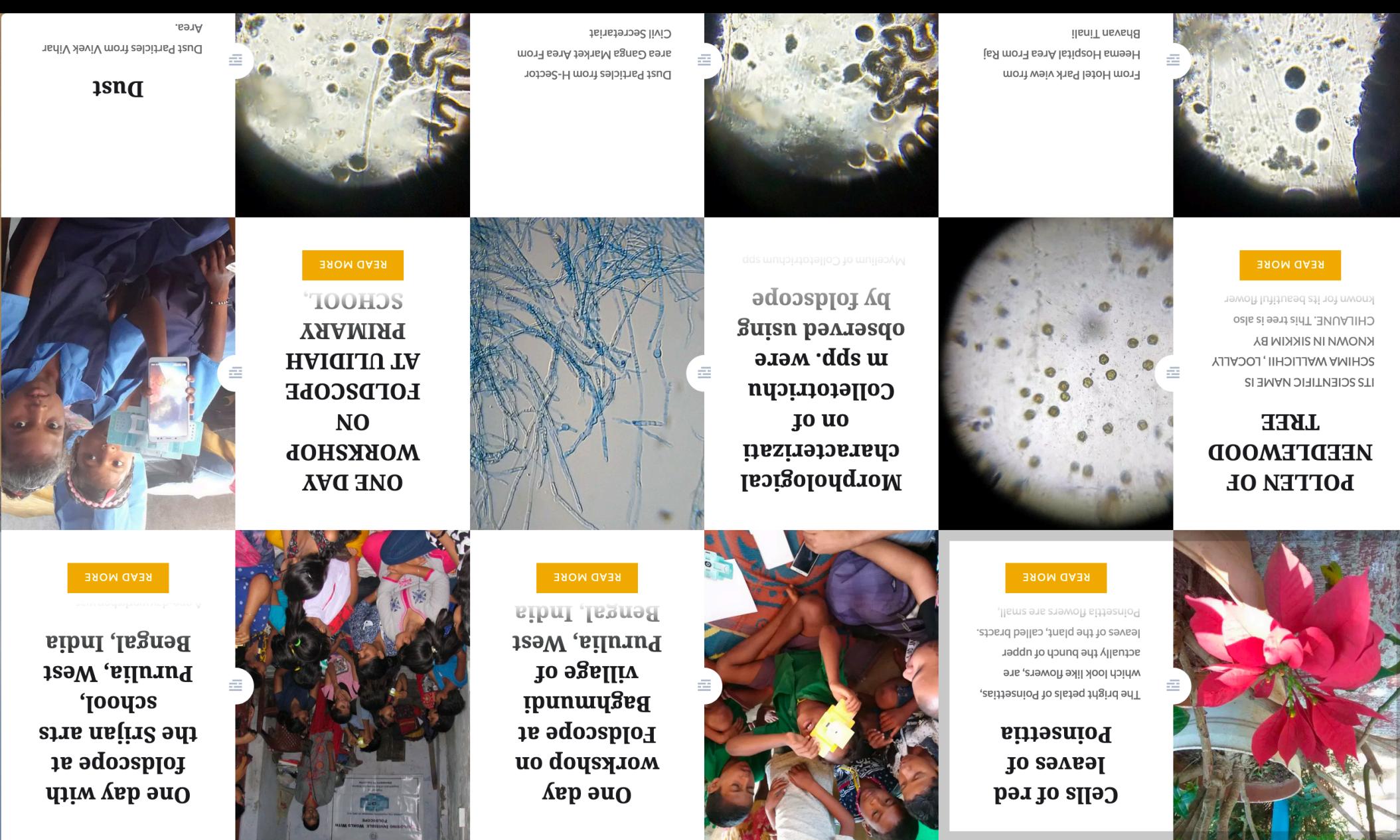
*Your project output must take the form of some digital artifact that can be shared on a world-readable basis. Video, article, recording, other

- Topic or aspect of your choosing
- Audience(s) of your choosing
- Method of your choosing*

Introduce Bioengineering

15% of grade

Intro. to BioE, Spring 2019
Final Project





Topic/Aspect Audience Method



Topic/Aspect Audience Method

4 Points

Teammate self assessments

30 Points

Picking topic/aspect, audience, method
(starting Week 6; 10 points each)

30 Points

Brainstorming topic/aspect, audience, method
Teams rule! (6 points)
(starting this Friday; 10 points each)

36 Points

Develop actual project output (NYT editorial, video, etc)
(starting Week 7, 10 points each for different aspects)

CLASS FRIDAY WILL BE HERE, IN 320-105

AND learn brainstorming...

Come to class this Friday to assemble into your teams

AND learn brainstorming...

Come to class this Friday to assemble into your teams

Step 2.

Team building quiz, right now...

Step 1.

FRIDAY CLASS WILL BE HERE, IN 320-105



FRIDAY CLASS WILL BE HERE, IN 320-105