



Customer Screening for DNA Synthesis

An interactive exercise from the International Biosecurity and Biosafety Initiative for Science



Instructions for workshop organizers

Slides 3-8 are intended as an introduction to synthesis and customer screening, to be presented in 5-10 minutes. You **may be able to skip some or all** of this depending on the background of your audience.

The Order Screening Game itself can be run in about 30 minutes:

- Slides 9-11 (<5 min) explain game, split into pairs, individuals read their profile
- Slide 12 (5 min) round 1 of screening
- Slides 13-14 (5 min) round 2 of screening game (roles reversed)
- Slide 15 (3 min) debrief in pairs
- Slide 16 (5 min) debrief in small groups
- Slide 17 time permitting, share back with the whole group

Slides 18-19 contain more information about IBBIS's mission and screening tools.

Want to run a workshop? Reach out to **screening@ibbis.bio** for assistance!



Instructions for workshop organizers

The latest version of the workshop materials (personas + instructions) can be found in:

Screening Game Slides (this slide deck; link will have the latest version) ibbis.bio/customer-screening-workshop-slides

Customer Profiles (formatted to print) ibbis.bio/customer-profiles-for-screening

An extra layer of organization is needed if you wish to run the game virtually or in a hybrid format. Please reach out to us for more specific guidance.

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- 1. Few people are motivated to do harm (good)
- 2. Doing harm with biology is difficult (good)
- **3.** ... because biology is difficult in general (bad)





Synthetic NAs are exponentially cheaper, while orders are longer and more complex

USD per base, 1999-2018. Data from Rob Carlson (2023) and Potomac (2018)

Plot generated in R using assistance from GPT4. Data from Carlson (2023) and Potomac (2018). With thanks to Max Langenkamp (see in-text links)

Custom orders ship in the mail, print on benchtops, or are sourced from third parties



Images: Founder's Pledge, Aric Crabb/Bay Area News Group, DNAScript



In 2006, a journalist orders smallpox DNA in the mail, spurring BWC talks and industry action

• This article is more than **19 years old**

Lax laws, virus DNA and potential for terror

• Loopholes mean anyone can order gene sequences • Scientists back voluntary regulation as first step

The smallpox virus last wreaked havoc on the human population in before a World Health Organisation programme eradicated it from the It now exists only in government laboratories in the US and Russia.

But ordering part of this long-dead pathogen's DNA proved easier the anyone dared imagine. All it took was an invented company name, a phone number, a free email address and a house in north London to the order by post.

What the investigation makes clear is that anyone, without any atter prove they are part of a legitimate research organisation, can order D sequences from any potential pathogen without fear of extensive questioning. In our case, VH Bio Ltd did not realise it was supplying the smallpox genome, but many scientists argue that it is the respon of companies selling custom-made pieces of DNA to check their orde potentially dangerous sequences.



Five Member Companies Represent 80 Percent of Worldwide Gene Synthesis Capacity

Washington D. C., Norember B, 2007–Free of the world's backing gover synthesis programs to day anomaly and an entropy of the program of the program of the programs of a synthesis of the program of the program of the program synthesis of synthesis of the program of the synthesis of program synthesis backs of the program of program of the synthesis of the program of program of the synthesis of the program of the program of the program of the global gover synthesis for the program of the program and others occurred to promote the beneficial application of gover probes is the program of t

The are private to attribution with diminatory of the intermediation of control synthesis in the synthesis of the synthesis of the synthesis of the synthesis is and Biokelmology. "The depth and breadth of expertise in gene synthesis represented by the participating companies, in concert with our dedication to policy based on sound science and thoughtful leadership, will enable us to shape the growth of a safe gene synthesis industry poieted to help address the technological needs of the 21^o century." **Today**, new standards respond to changing risk landscape (AI tools, long synthesis, biofoundries)





Acquisition: reconstructing a functional pathogen or toxin from digital sequences

How Canadian researchers reconstituted an extinct poxvirus for \$100,000 using mail-order DNA

A study that brought horsepox back to life is triggering a new debate about the risks and power of synthetic biology —

6 JUL 2017 · BY KAI KUPFERSCHMIDT



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Article | Published: 04 May 2020

Rapid reconstruction of SARS-CoV-2 using a synthetic genomics platform

Tran Thi Nhu Thao, Fabien Labroussa, Nadine Ebert, Philip Vžkovski, Hanspeter Slaider, Jasmine Portmann, Jenna Kelly, Silvio Steiner, Melle Holwerda, Annika Kratzel, Mitra Gultom, Kimberly Schmied, Laura Laloli, Linda Hüsser, Manon Wider, Stephanie Pfaender, Dany, Hirt, Valentina Ciopab, Silvia Crespo-Pomar, Simon Ströder, Dorzeen Muth, Daniela Niemeyer, Victor M. Corman, Marcel A. Müller, ... Volker Thiel 🖾 + show authors

Nature 582, 561–565 (2020) | Cite this article

151k Accesses | 256 Citations | 1506 Altmetric | Metrics

Engineering: enabling more people to engineer pathogens and toxins

1. Recognize potentially risky sequences toxins, pathogen genomes, virulence factors

2. Decide whether to trust user or customer with risky sequences by screening legitimacy

1. Recognize potentially risky sequences toxins, pathogen genomes, virulence factors

2. Decide whether to trust user or customer with risky sequences by screening legitimacy

Legitimate customers come in many shapes and institutions

- Orders from academic researchers could be centralised across entire departments or conducted by individual undergraduates
- Early-stage **biotechnology companies** may lack legal standing, lab space, online presence, and other indicators of legitimacy
- Third parties like **CROs**, **CMOs**, **and biofoundries** may not have designed the experiments for which they order DNA
- Biosafety best practices **vary around the world**, including whether formal review is conducted

Order Screening Game

An interactive exercise from the International Biosecurity and Biosafety Initiative for Science

Each of you will play a synthesis screener and a customer.

Customer profiles based on real examples of legitimate scientists and attempted bioterror

Flagged orders screened using IBBIS's Common Mechanism

Step one: who are you?

- 1. Read your customer profile.
- 2. Find a partner. **Do not show** them your profile!
- 3. Decide who will play the customer first. The other person will be the screener.
- 4. The customer should hand their printed order to the screener.

As the customer, you want the screener to send the sequence.

As the screener, you must decide to:

- 1. Fulfill the order
- 2. Deny the order
- 3. Deny and <u>report to law</u> <u>enforcement</u>

switch!

without sharing your true identity, change screener and customer roles.

As the customer, you want the screener to send the sequence.

As the screener, you must decide to:

- 1. Fulfill the order
- 2. Deny the order
- 3. Deny and <u>report to law</u> <u>enforcement</u>

reveal!

Show your partner your customer profile.

Take 3 minutes to discuss if this is surprising, and if you would change your decision about whether to fulfil the order.

Join 1-2 other pairs.

For 5 minutes, discuss:

- What did you notice?
- What information did you wish you had?
- What systems would help with follow-up screening?

Share a question or observation from your group with the rest of the room.

You can use IBBIS screening tools right now

com/mec

commec: a free, open-source, globally available tool for sequence screening

customer screening forms and decision guidance to help rigorously verify legitimacy

Reach out to **screening@ibbis.bio** with questions or for help getting started!

Safeguarding modern bioscience and biotechnology so it can advance and flourish safely and responsibly

for DNA synthesis:

Increasing the share of synthesis orders for which sequences and customers are screened

Supporting international standards that are inclusive and rigorous