

Screening of microbiome of bark beetle *Ips typographus* for antibiotic producers

Sofiia Sinelnikova^{1,2}, Anna A. Baranova¹, Vera A. Alferova¹, Olga A. Belozeroва¹, Arseniy A. Sinichich^{1,3}

¹ Shemyakin-Ovchinnikov Institute of Bioorganic Chemistry, Miklukho-Maklaya 16/10, 117997 Moscow, Russia

² Department of Biotechnology, A.P. Nelyubin Institute of Pharmacy, I.M. Sechenov First Moscow State Medical University (Sechenov University), Trubetskaya 8, building 2, Moscow, 119991, Russia

³ Department of Chemistry, Lomonosov Moscow State University, Leninskie Gory 1, Moscow 119991, Russia

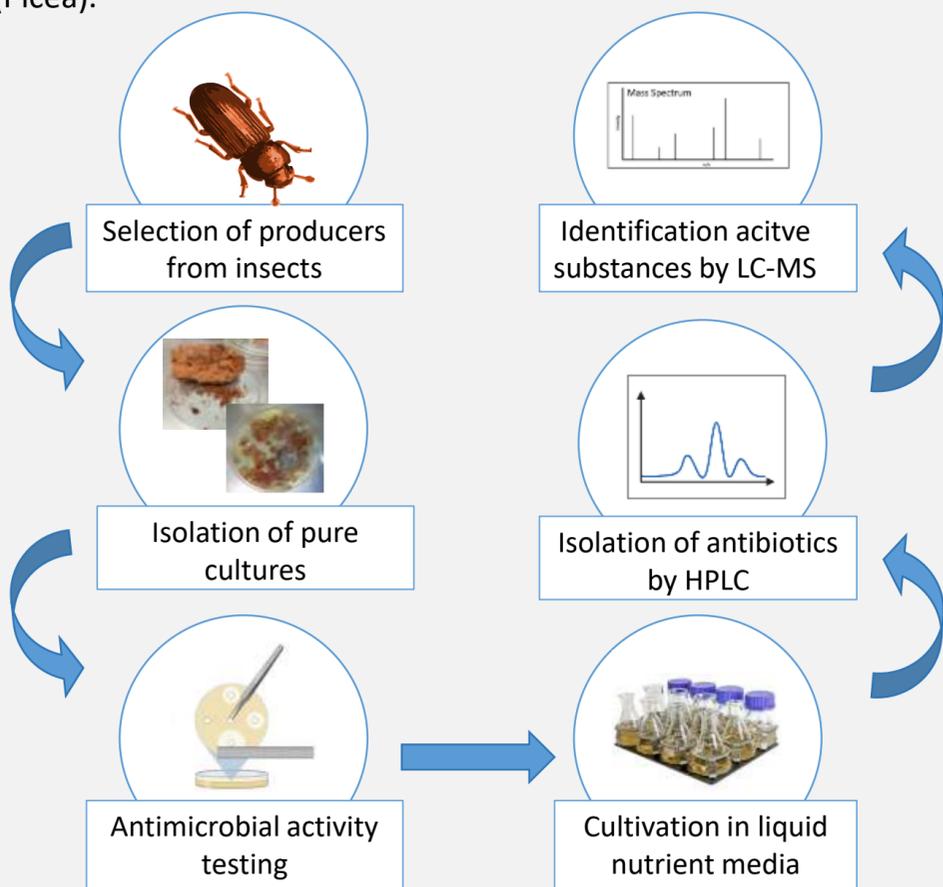
email: sinelnikowa.sofia@yandex.ru

INTRODUCTION

One way to search for antibiotics is the isolation of microbial antibiotic producers from insects and their products. Symbionts and insect-associated microorganisms represent an almost inexhaustible source of bioactive compounds, including antibiotics

OBJECT OF STUDY

In this study, we examined samples of the bark beetle *Ips typographus* found in the bark of fallen spruce (*Picea*).



Flowchart of the research process

ANTAGONISTIC AKTIVITY

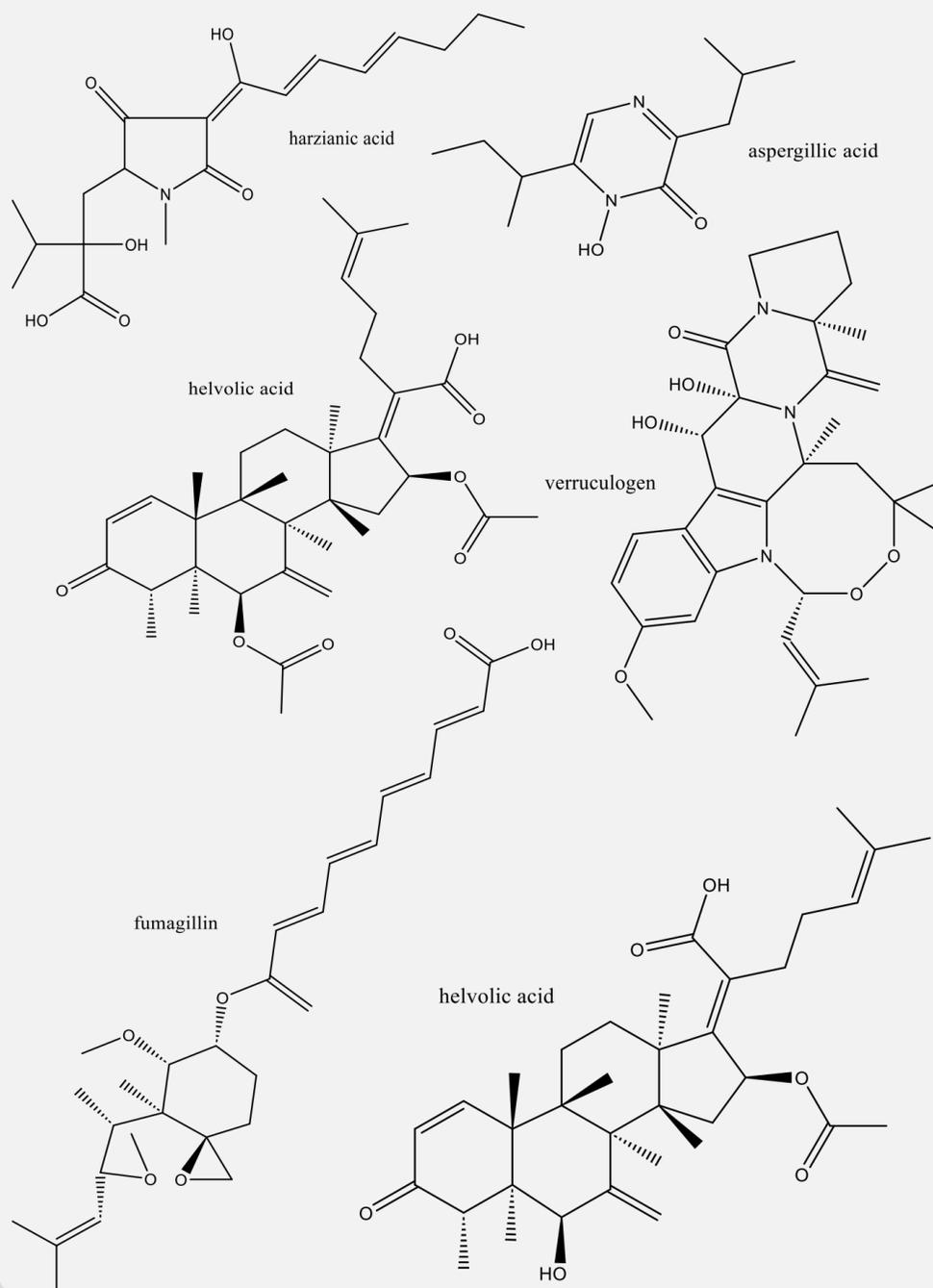
113 microbial isolates were obtained. Antimicrobial activity was investigated using the agar diffusion method against a wide range of tested microorganisms. The table includes 16 isolates with any activity indicators.

| Ips typographus | <i>A. niger</i> INA 00760 | <i>C. albicans</i> CBS 8837 | <i>B. subtilis</i> ATCC 6633 | <i>E. coli</i> ATCC 25922 | <i>E. faecalis</i> ATCC 29212 | <i>E. coli</i> dtolC | <i>E. coli</i> lptD |
|-----------------|---------------------------|-----------------------------|------------------------------|---------------------------|-------------------------------|----------------------|---------------------|
| K1- | 26 | + | - | +++ | - | - | ++ |
| | 27 | +++ | - | + | - | - | + |
| | 31 | - | - | - | - | - | + |
| K2- | 6 | - | ++ | ++ | - | + | ++ |
| | 7 | - | ++ | ++ | - | + | ++ |
| | 20 | - | ++ | ++ | - | - | ++ |
| K3- | 12 | - | - | + | - | - | ++ |
| | 13 | - | - | + | - | - | + |
| K4- | 10 | - | + | ++ | - | ++ | + |
| | 14 | - | - | - | - | - | + |
| | 15 | - | - | - | - | - | + |
| | 27 | - | - | + | - | - | ++ |
| | 28 | - | - | + | - | - | +++ |
| | 30 | - | - | + | - | - | + |
| | 31 | - | - | - | - | - | + |
| | 33 | - | - | - | - | - | ++ |

«+++» - high activity, «++» - moderate activity, «+» - low activity, «-» no activity

ISOLATION OF ANTIBIOTICS

As a result, 3 isolates of micromycetes with pronounced antimicrobial properties were selected for further study and cultivation in liquid nutrient media. The active substances were identified by LC-MS. Strain K1-26 produced harzianic acid, which has antifungal activity and is an inhibitor of acetoxyacid synthase. Strain K4-28 produced aspergillilic acid, which is also known as an antibiotic with antifungal activity. Strain K2-6 produced several antibiotic substances: verruculogen (diketopiperazine alkaloid with broad spectrum of biological activity), fumagillin (mycotoxin), helvolic acid (a broad-spectrum mycotoxin), helvolinic acid (showed potent antimicrobial activities against *Staphylococcus aureus*).



CONCLUSIONS

- Screening the *Ips typographus* microbiome revealed three micromycete isolates with significant antimicrobial activity
- LC-MS analysis identified the production of several known antibiotics, suggesting the bark beetle microbiome as a valuable source for novel bioactive compounds.