جامعة الملك عبدالله للعلوم والتقنية King Abdullah University of Science and Technology

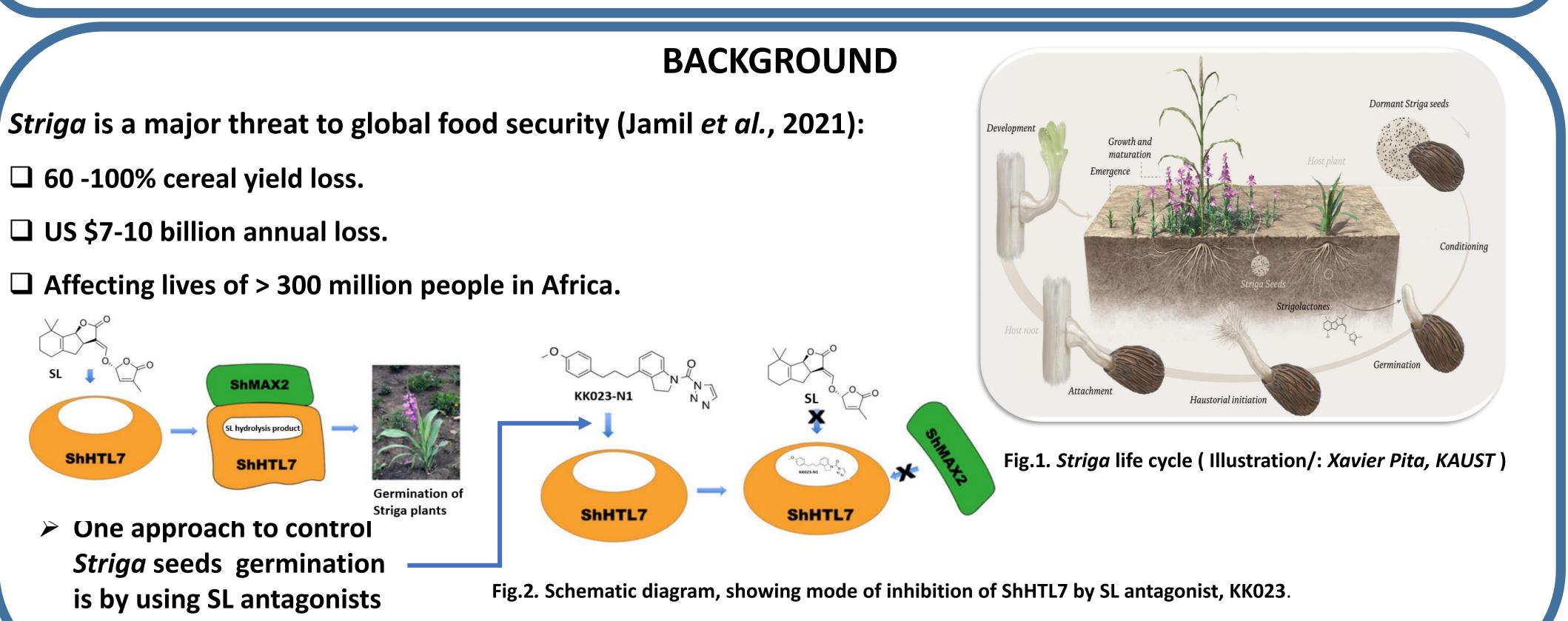
# <u>Randa Zarban<sup>1</sup></u>, Muhammad Jamil<sup>1</sup>, Umar Hameed<sup>1</sup>, Jian You Wang<sup>1</sup>, Tadao Asami<sup>2</sup>, Stefan Arold<sup>1</sup>, and Salim Al-Babili<sup>1\*</sup>.

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## SUMMARY

- Striga is a root parasite, attacking staple cereal crops with huge yield losses
- High production of long lived tiny *Striga* seeds, with complex life cycle has made *Striga* control very difficult.
- We have developed a very promising *Striga* specific inhibitor "KK023".
- KK023 inhibit *Striga* seed germination by blocking ShHTL7 receptor.
- This compound will lead to the development of inhibitors to overcome *Striga* infestation, a threat to global food security.

### □ Affecting lives of > 300 million people in Africa.



Synthesis of KK023:

(Nakamura *et al.,* 2019)

(Hameed *et al.,* 2019)

In vitro YLG hydrolysis assay and In silico modeling: YLG hydrolysis by ShHTL7±KK023 was carried out in 96-well black plate and fluorescence was detected by SpectraMax i3 device. Modeling of ShHTL7 binding to KK023 was done using SwissDock server (Zarban et al., 2021).

Striga and rice germination:

Germination of Striga seeds was evaluated in response to GR24±KK023, Triton X-100, or KK094. Rice seeds (*cv* Nipponbare), were grown hydroponically  $\pm$  KK023, TritonX-100, or KK094 (Zarban *et al.*, 2021).

Striga emergence (greenhouse study):

Rice (*cv* IAC-165) plants were grown in *Striga* infested soil and were treated with 10 or 100 μM of KK023 or Triton X-100 for four weeks (twice a week) (Zarban *et al.*, 2021). *Striga* emergence in each pot was observed.

# Designing specific seed germination inhibitors to combat Striga for tackling food security problem in Africa

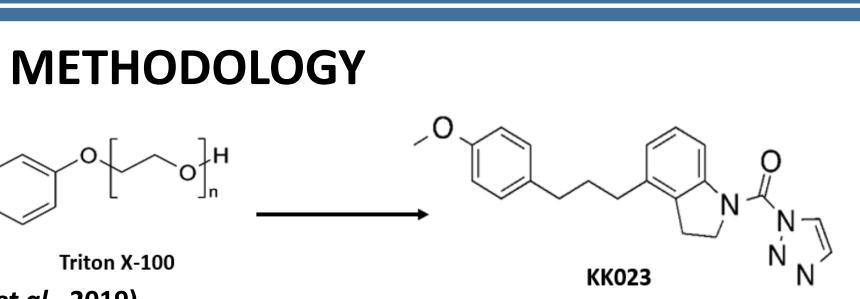
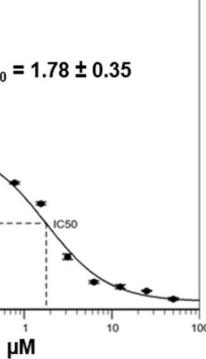
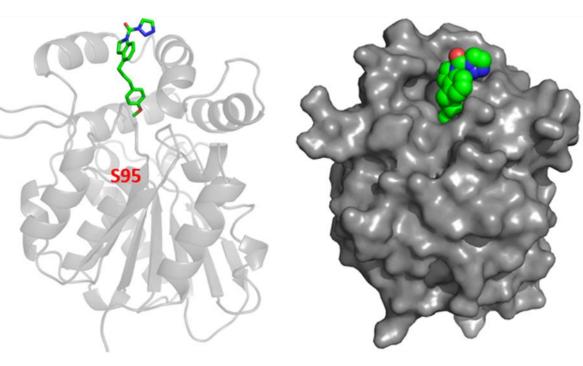


Fig.3. Synthesis and chemical structure of KK023 inhibitor

# **1. KK023 inhibited ShHTL7 mediated- YLG hydrolysis:** ShHTL7/KK023-N1 $IC_{50} = 1.78 \pm 0.35$ Fig.4. KK023 inhibited ShHTL7 enzymatic activity by binding non-covalently to ShHTL7 active pocket residues (model: ShHTL7 (grey), KK023 (green)). 2. KK023 inhibited *Striga* seeds germination: ■ KK023-N2 □ KK023-N1 □ KK094 ত 50 ൾ 10 10 Co-application (uM) with GR24 (1.0nM Fig.5. Application of KK023, Triton X-100 and KK094 inhibited the germination of Striga seeds. less negative impact on host crop, unlike KK094 or Triton X-100. Saharan Africa. Funding This study has been financially supported by the Bill & Melinda Gates Foundation grant OPP1194472 and

RESULTS





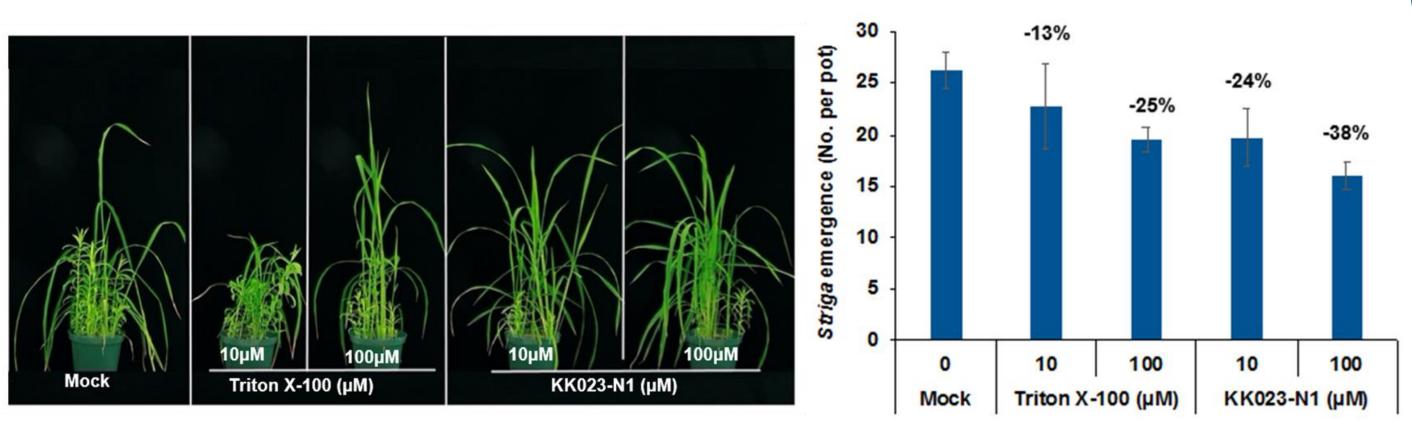
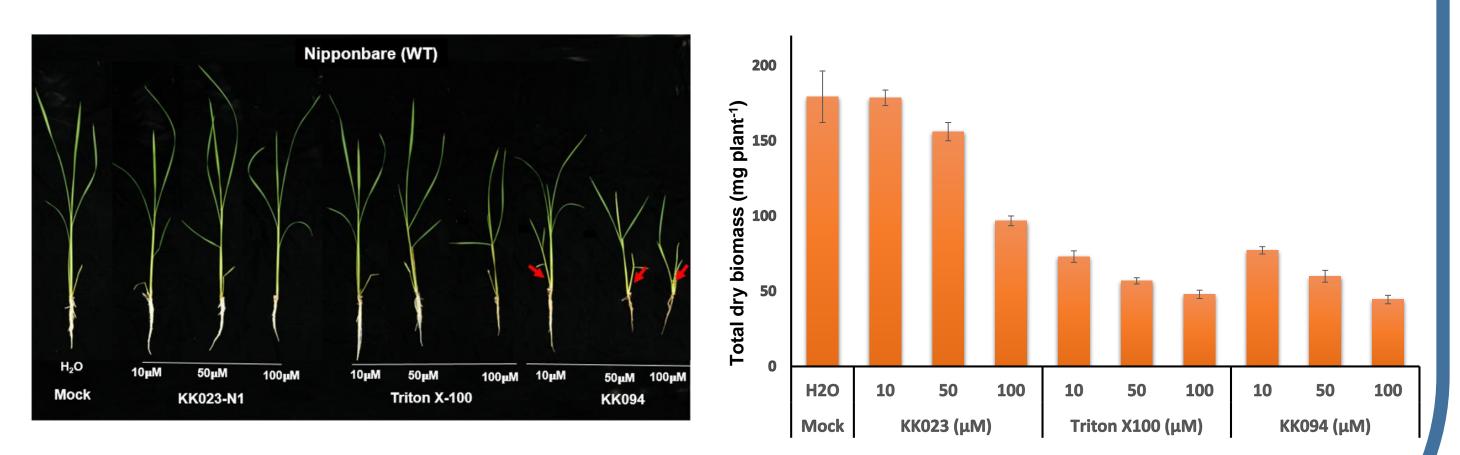


Fig.6. Application of KK023 and Triton X-100 reduced Striga emergence under greenhouse conditions, however, KK023-N1 caused 38% reduction with less negative impact on rice as a host crop.

### 4. KK023 is save and imposing no effect on host SL signaling or growth:



## CONCULSION

**KK023** inhibitors are novel chemicals which specifically bound to ShHTL7, reduced the Striga seeds germination and emergence of Striga plants, with

**KK023** can be a promising herbicide to combat Striga seed germination and infestation for tackling global food security issues, specifically in sub-

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### 3. KK023 is a promising inhibitor to combat *Striga* infestation

Fig.7. KK023 has no effect on SL signaling or enhance the emergence of the second tiller as KK094 and did not reduce the biomass of rice as a host crop.

