Residue dissipation, degradation dynamics, and dietary risk assessment of mandipropamid in ginseng under field application

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1 **Residue dissipation, degradation dynamics, and dietary risk assessment of**
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3 **Running title: Degradation and assessment of mandipropamid in ginseng**

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10 Abbreviations

11

Abstract: Mandipropamid, a new fungicide for oomycete disease, has a strong effect on the blight of many crops and has been registered for the treatment of ginseng blight in China. However, no maximum residue limit (MRL) of mandipropamid has been identified for ginseng, and there have been few related studies. We established and verified the analysis method of mandipropamid in ginseng using high-performance liquid chromatography-tandem mass spectrometry. The method has good linearity and accuracy in the range of 0.002–0.5 mg/kg. The average recovery of mandipropamid was 87.4–101.6%, and the standard deviation was 1.1–10.1. Mandipropamid in ginseng plants and soil rapidly degraded following first-order kinetics models. The degradation dynamics showed that the half-life of mandipropamid in ginseng plant and soil was 13.8–28.0 and 9.8–27.4 d, respectively. After the recommended dose of mandipropamid was applied once, the residual content of mandipropamid in fresh ginseng, dried ginseng, red ginseng, ginseng plant, and ginseng soil was <0.01–0.185, <0.01–0.265, 0.085–1.544, 0.075–4.800, and <0.01–0.014 mg/kg, respectively. The dietary risk assessment of mandipropamid on ginseng showed that the risk quotient value was far less than 100%, indicating that the recommended dose of mandipropamid does not cause unacceptable risks to humans. After the recommended dose of mandipropamid was applied once, it did not cause unacceptable risks to humans. This study not only provides a reasonable spray dosage of mandipropamid to ginseng, but also offers a reference for the establishment of MRLs in China.

32 **Keywords:** mandipropamid; ginseng; HPLC-MS/MS; dissipation; dietary risk

33 assessment

34

1 Introduction

Ginseng is the dried root and rhizome of *Panax ginseng* C.A.Mey (Zhang et al., 2021). As the largest ginseng planting and production country in the world, China has a long history of medicinal use of ginseng. Ginseng plays an important role in regulating human health and treating different diseases (Fan et al., 2020) through its antitumor, antioxidation, and antidiabetic properties (Park et al., 2017). The growth cycle of ginseng is relatively long, usually 4–6 years for garden ginseng and more than 15 years for forest ginseng. During the long growth cycle, the occurrence of diseases affects the quality of ginseng, which causes significant economic losses to ginseng farmers and affects the import and export of ginseng. Therefore, fungicides must be used to prevent diseases during the planting process. However, due to the irregular use of pesticides, excessive pesticide residues can be found in ginseng.

Mandipropamid is a new fungicide for oomycete diseases (Tang et al., 2011), which is the first commercial amygdalinamide compound with the chemical name of 2-(4-chlorophenyl)-n-[2-(3-methoxy-4-prop-2-ynynloxy-phenyl)-ethyl]-2-prop-2-ynyloxy-acetamide. It has high activity in inhibiting conidial germination, mycelial growth, and spore formation; It also has a good preventive and therapeutic effect on lychee downy mildew (Tang et al., 2011), pepper blight (Siegenthaler and Hansen, 2021), tomato late blight (Chen et al., 2021), and cucumber downy mildew (Liu et al., 2016; Zhang et al., 2015). In China, mandipropamid has been registered in tomato, pepper, potato, lychee, grape, watermelon, and ginseng and has been widely used to control

56 blight and downy mildew in various crop sites. Mandipropamid is mainly used to treat
57 ginseng blight. The application of mandipropamid to ginseng will inevitably cause
58 pesticide residue problems in ginseng and the surrounding environment, and the 2008
59 JMPR established an ADI for mandipropamid of 0-0.2 mg/kg bw/day. It is worth
60 noting the widespread use of mandipropamid whether will have any impact on human
61 health. And in recent years, the problem of pesticide residue has gained increasing
62 attention; therefore, the rational use and risk assessment of mandipropamid is of great
63 significance.

64 At present, the methods for detecting residues of mandipropamid in sesame leaves
65 (Farha et al., 2016), grapes (Kwon et al., 2008; Malhat and Abdallah, 2012; Xu et al.,
66 2020; Zhang et al., 2014), tomato (Kwon et al., 2008), pepper (Kwon et al., 2008;
67 Zhang et al., 2014), cabbage (Kwon et al., 2008), potato (Kwon et al., 2008; Zhang et
68 al., 2014), watermelon (Zhang et al., 2014), lettuce (Yang et al., 2020), and *Cnidium*
69 (Dong et al., 2020) are mainly high-performance liquid chromatography (HPLC) and
70 HPCL-tandem mass spectrometry (MS/MS). Moreover, residual behaviour analysis
71 and research on mandipropamid in grapes (Malhat and Abdallah, 2012), lettuce (Yang
72 et al., 2020), and *Cnidium* (Dong et al., 2020) have been undertaken. However, there
73 have been no systematic studies on the residual behaviour, degradation dynamics, and
74 dietary risk assessment of mandipropamid in ginseng. Therefore, in this study, we
75 established a method for the detection of mandipropamid in ginseng using HPLC-
76 MS/MS. Through field trials at five locations for 1 year, the residual behaviour and

77 degradation dynamics of mandipropamid in ginseng were analysed. Additionally, the
78 safety of mandipropamid use on ginseng was evaluated in combination with the
79 Chinese dietary structure. Finally, reasonable suggestions were provided for the
80 rational use of mandipropamid on ginseng, and the basis for the formulation of
81 residue limits in ginseng was provided.

82 **2 Materials and Methods**

83 **2.1 Instruments and reagents**

84 The following materials were used: 1260-6470 HPLC-MS/MS (Agilent
85 Technologies, USA), SHMADIU Model AUY220 Balance (Shmadiu Corporation,
86 Japan), HUAPUDA Model YXJ-A Centrifugal Machines (Changzhou Huapuda
87 Educational Instrument Corporation, China), Kunshan KQ-500 Ultrasonicator
88 (Kunshan Ultrasonics Corporation, China), and a digital high-speed dispersive
89 homogeniser (IKA T25, IKA Company, Germany).

90 Analytical standards of mandipropamid (purity 99.5%, Dr. Ehrenstorfer GmbH,
91 Germany) were used. The mandipropamid standard substance (0.0101 g) was
92 accurately weighed, diluted to 10 mL with acetonitrile, sonicated for 10 min, and
93 stored in a refrigerator at -4°C . Acetonitrile and methanol were of HPLC grade
94 (MREDA Technology Inc., USA), formic acid was of 96% HPLC grade (TEDIA
95 Company, USA), ammonium acetate was of 98% analysis grade (SIGMA, Germany),
96 both sodium chloride and magnesium sulphate were of analytical grade (Beijing

97 Chemical Plant, China), and Cleanert PSA was of PR grade (Agela Technology,
98 China).

99 **2.2 Field trial**

100 **2.2.1 The degradation experiments of mandipropamid in ginseng plant and soil**

101 The degradation experiments of mandipropamid on ginseng plants and soil were
102 carried out in Baishan City, Jilin Province, Huanren Manchu Autonomous County,
103 Liaoning Province, and Fusong County, Jilin Province, respectively. One plant
104 degradation experiment plot and one soil degradation experiment plot were set up,
105 each plot was 50 m², and repeated three times. In the early stage of ginseng blight, the
106 dosage of 900 mL/ha (210.6 g a.i./ha) was applied once, and the water was mixed
107 with 600 kg/ha. Ginseng plant and soil samples were collected at 0, 1, 3, 5, 7, 14, 21,
108 28, 35, 45, and 60 days after application.

109 **2.2.2 The final residue experiments of mandipropamid in ginseng**

110 According to (Pesticide Residue Test Guidelines) (Ministry of Agriculture and
111 Rural Affairs of the People's Republic of China, 2018), the final residue experiments
112 were carried out in Yanji City, Jilin Province, Baishan City, Jilin Province, Huanren
113 Manchu Autonomous County, Liaoning Province, Fusong City, Jilin Province, and
114 Ji'an City, Jilin Province. Three experiments plots were set up, each plot was 50 m².
115 In the early stage of ginseng blight, the dosage of 900 mL/ha (210.6 g a.i./ha) was

applied once, and the water was mixed with 600 kg/ha. Fresh ginseng, ginseng plant and soil samples were collected twice at 14, 21, and 28 days after application.

2.2.3 Sample collection

Collection of fresh ginsengs: 2 kg of normal and disease-free underground fleshy root of ginseng was randomly collected with 12 plots, and the soil attached to ginseng was brushed off with a brush and put into a plastic sealing bag.

Collection of ginseng plants: 1 kg of normal and disease-free ginseng aboveground parts were randomly collected with 12 plots, cut into 1 cm size with scissors, and mixed evenly. 500 g of samples were taken by four methods and put into plastic sealing bags.

Collection of ginseng soil: a soil sampler was used to randomly collect ginseng soil samples at five points, and 0-10 cm deep soil was collected in the degradation experiments, and 0-15 cm deep soil was collected in the final residue experiment. After the collected soil was thoroughly mixed, the impurities such as gravel and sticks were removed and 500 g samples were taken by quartering method into a plastic sealing bag.

2.2.4 Preparation of samples

Fresh ginseng samples were gently washed using cool water, and the surfaces of the samples were dried indoors. Fresh ginseng samples were baked at 55°C for 24 h to prepare dried ginseng, and fresh ginseng samples were placed in a cage on a pot filled

with water at 80°C and heated to boiling for 2.5–3 h. Then, the heated ginseng was cooled and dried at 70°C for 6 h with a dryer. Next, the ginseng was infiltrated with water, and the ginseng was dried again at 55°C for 24 h. Ginseng plant samples were placed in a tissue masher, and mashed for 5 min.

2.3 Extraction and purification

Next, 5.0 (± 0.05) g of the homogenised fresh ginseng, dried ginseng, red ginseng, ginseng plant, and ginseng soil samples were added to 50-mL centrifuge tube, and then, 5 mL of distilled water and 10 mL of acetonitrile were added to the tube. After standing immersion for 5 min, vortex extraction was performed for 2 min, and the samples were centrifuged at 5,000 g for 5 min.

An aliquot of 1 mL of the upper layer of fresh ginseng, dried ginseng, red ginseng, and ginseng soil was transferred to a centrifuge tube containing 50 mg PSA and 100 mg anhydrous magnesium sulphate. An aliquot of 1 mL of the upper layer of the ginseng plant was transferred to a centrifuge tube containing 50 mg PSA, 25 mg GCB, and 100 mg anhydrous magnesium sulphate. The mixture was shaken vigorously 100 times and centrifuged for 5 min at 5,000 rpm. The upper layer was filtered through a 0.22- μ m nylon syringe filter disc and analysed by HPLC-MS/MS.

2.4 Instrumental parameters

2.4.1 Chromatographic conditions

155 The chromatographic column used was an Agilent Zorbax RRHD Eclipse Plus C18
156 (3.0 ×100 mm, 1.8 µm at 30°C) column. The mobile phase was composed of
157 acetonitrile (A) and water containing 0.1% formic acid and 5 mmol ammonium
158 acetate (B) with a linear gradient. The gradient elution procedure is shown in Table
159 S1.

160 **2.4.2 Mass spectrometry conditions**

161 The ionisation source mode was electrospray ionisation, ionisation source polarity
162 was positive ion mode (ESI+), scanning mode was multiple reaction monitoring,
163 atomising gas was nitrogen, nebuliser gas pressure was 45 psi, ion spray voltage was
164 4,000 V, sheath gas temperature was 350°C, sheath gas was 12 L/min, quantitative
165 ion of mandipropamid was 412/328.1, qualitative ion of mandipropamid was
166 412/328.1 and 412/124.8, fragmentor was 60 V, and collision energy was 412/328.1
167 (13 V) and 412/124.8 (35 V).

168 **2.5 Configuration of standard solutions**

169 Accurately weighing 0.0101 g of mandipropamid standard, using acetonitrile
170 constant volume to 10 mL, prepared with a concentration of 1000 mg/kg of standard
171 mother liquor, stored in a refrigerator at -18°C, shelf life of 6 months. During use,
172 acetonitrile and blank matrix solution were diluted to 0.001, 0.002, 0.005, 0.01, 0.02,
173 0.05, 0.1, 0.2, 0.5 mg/kg, respectively. According to the above instrument conditions,

the injection concentration as horizontal coordinate, linear regression model was established with the peak area as ordinate.

2.6 Statistical analysis

In this study, the standard curve and matrix standard curve of the same concentration were detected under the same chromatographic conditions, and the matrix effect (ME) was calculated according to the slope of the two standard solutions. When the matrix effect was within 100% (± 20), it was ignored; when it was greater than 120%, it showed enhancement, and when this was less than 80%, it showed inhibition. The calculation formula is as follows (Fang et al., 2019):

$$ME(\%) = K_s / K_m \times 100(1)$$

where K_s is the slope of the standard curve and K_m is the slope of the matrix standard curve.

The dissipation dynamics of mandipropamid in ginseng plants and soil were evaluated using first-order kinetics as follows (Fang et al., 2019; Vargas-Pérez et al., 2020):

$$C_t = C_0 \times e^{-Kt}(2)$$

$$T_{1/2} = \ln 2 / k(3)$$

where C_t (mg/kg) is the residual concentration of mandipropamid in the ginseng plant or soil, C_0 (mg/kg) is the initial concentration of mandipropamid in the ginseng plant or soil, k (day^{-1}) is the dissipation rate constant, and $T_{1/2}$ is the pesticide half-life of

pesticide degradation.

2.7 Dietary intake risk assessment

Dietary exposure and risk assessments were performed to ensure the rational use of mandipropamid. (Risk Quotient) $RQ > 100\%$ indicated that there was an unacceptable risk to human health; the larger the value, the greater the risk. $RQ < 100\%$ indicated that the risk to human health was acceptable; the smaller the value, the smaller the risk.

The chronic dietary risk assessment was undertaken by calculating RQ as follows (Fang et al., 2019):

$$NEDI = (\sum STM Ri \times Fi) / bw(4)$$

$$RQ = NEDI / ADI \times 100 \quad (5)$$

where NEDI (mg/kg, bw) is the national estimated daily intake, STM Ri (mg/kg) is the supervised trial median residue level, Fi (kg/d) is the average daily food intake, bw (kg) is the average Chinese body weight (63 kg), and ADI (mg/kg, bw) is the acceptable daily intake.

3 Results and Discussion

3.1 The Optimization of extraction and purification

In this study, the optimization of extraction and purification were conducted before sample testing. Different volumes (10, 15 and 20 mL) of Acetonitrile, Methanol,

Ethyl acetate and Acetone were selected for comparison. The result shows the extraction effect of 10 mL of acetonitrile can meet the recovery requirements and reduce the amount of solvent used. Thus, 10 mL 10 mL of acetonitrile was chosen as the extraction solvent. The most commonly used scavengers for the QuEChERS method are PSA, C18, GCB, and anhydrous MgSO₄. We compared the purification effects of different doses of PSA, C18 and GCB and found that 50 mg PSA and 100 mg anhydrous magnesium sulphate has good purification effect for all samples. Due to the ginseng plant containing a lot of pigment, add 25 mg GCB as scavengers to clean up ginseng plant sample.

3.2 Method validation

The standard solution was then diluted with acetonitrile and blank matrix solution to 0.002, 0.005, 0.01, 0.02, 0.05, 0.1, 0.2, and 0.5 mg/L series solutions. Instruments detection was performed according to the 2.3 method. With the concentration as the abscissa and the peak area as the vertical coordinates, a standard curve was drawn, resulting in the linear regression equation of mandipropamid. The results are presented in Table 1, the correlation coefficients were all greater than 0.99. The ME of fresh ginseng, dried ginseng, ginseng plant and soil were within 100 ($\pm 20\%$), indicating that the samples of them can be quantified by standard curve, and the ME of red ginseng was 61.2%, which indicate that red ginseng shown the inhibition for mandipropamid, thus, the samples of red ginseng were quantified by matrix standard curve of red ginseng.

Table 1 The linear regression equation of mandipropamid in different matrices

Matrix	Linear regression equation	Coefficient (R)	ME (%)
Solvent	$y = 103408618.17 x + 1955282.89$	0.9854	/
Fresh ginseng	$y = 57171927.30 x + 199740.39$	0.9999	87.0
Dried ginseng	$y = 66912235.11 x + 20,781.54$	0.9996	91.7
Red ginseng	$y = 3967045.54 x - 588.30$	0.9999	61.2
Ginseng plant	$y = 101461225.21 x + 1330231.11$	0.9912	98.1
Ginseng soil	$y = 85006757.21 x + 653659.39$	0.9967	99.2

Note: ME, Matrix effects.

The accuracy and precision of the method were verified using an addition-recovery test. According to the above method, the added amounts of fresh ginseng and soil were 0.01, 0.05, and 0.5 mg/kg, and those of dry ginseng, red ginseng, and plants were 0.01, 0.05, 0.5, and 50 mg/kg, respectively. This method was repeated five times. The results are presented in Table 2. The sensitivity and accuracy of the method met the requirements and were suitable for the residue analysis of mandipropamid in ginseng.

247 Table 2 The addition recovery rate, RSD, LOQ, and LOD in ginseng

Matrix	Spiked level (mg/kg)	Average Recovery (%)	RSD (%)	LOQ (mg/kg)	LOD (mg/kg)
Fresh ginseng	0.01	89.9	10.1		
	0.05	87.4	4.0		4.3×10 ⁻⁴
	0.5	96.3	1.7		
Dried ginseng	0.01	96.1	1.3		
	0.05	99.0	1.8		
	0.5	101.6	2.8		3.6×10 ⁻⁴
	50	95.7	1.4		
Red ginseng	0.01	92.6	3.6		
	0.05	98.7	3.1		
	0.5	100.0	1.4	0.01	4.8×10 ⁻⁴
	50	100.1	2.7		
Ginseng plant	0.01	89.6	2.4		
	0.05	99.1	2.4		
	0.5	94.2	1.9		4.1×10 ⁻⁴
	50	95.8	2.2		
Ginseng soil	0.01	97.3	1.1		
	0.05	93.7	1.2		3.9×10 ⁻⁴
	0.5	96.8	1.5		

248 Note: RSD, relative standard deviation; LOQ, limit of quantitation; LOD, limit of
 249 etection.

250 3.3 The degradation dynamics of mandipropamid in ginseng plant and soil

251 The samples of ginseng plant and soil were detected using the 2.3-2.4 methods, the
 252 degradation of mandipropamid conformed to first-order kinetics. The content of

mandipropamid in ginseng plant and soil 2 h after application was that of the initial residues, and as time passed, the residue concentration of mandipropamid gradually decreased. The results are shown in Figure 1–2 and Table 3. The half-life of mandipropamid in ginseng plant was 13.8 (Fusong), 28.0 (Huanren), and 13.9 d (Baishan). The initial residue amounts of mandipropamid in ginseng plant were 5.24 (Fusong), 3.22 (Huanren), and 4.42 mg/kg (Baishan). The half-life of mandipropamid in ginseng plant was 9.8 (Fusong), 16.0 (Huanren), and 27.4 d (Baishan). The initial residue amount of mandipropamid in ginseng plants was 0.039 (Fusong), 0.054 (Huanren), and 0.068 mg/kg (Baishan). The difference in the initial residues may be due to multiple factors, such as plant growth status and weather conditions (temperature, humidity, wind speed, etc.) at the test point during spraying. The results showed that the half-life of mandipropamid in ginseng plants was not significantly different in Fusong and Baishan, whereas that in Huanren was significantly higher than that in Fusong and Baishan, mainly due to plant size, sparseness, and metabolic capacity, which may also be affected by factors such as plant variety and weather conditions (Bai et al., 2020). The half-life of mandipropamid in ginseng soil was different in Fusong, Huanren, and Baishan. Since before ginseng planting, the soil was artificially mixed with some other substances such as sand, its organic matter content varied between locations. The organic acid content of Baishan soil was 4%, that of Fusong soil was 4.3%, that of Huanren soil was 4.4%, and Baishan soil had the lowest organic matter content. This may be responsible for the half-life of

274 mandipropamid in Baishan soil, which was significantly higher than that of the other
 275 two locations, possibly because of to the effects of factors such as the types of
 276 microorganisms in the soil, temperature and humidity of the environment, and
 277 duration of sunshine.

278

279 Figure 1 The degradation dynamics curve of mandipropamid in ginseng plant

Figure 2 The degradation dynamics curve of mandipropamid in ginseng soil

Table 3 The degradation dynamics of mandipropamid in ginseng plant and soil

Matrix	Location	Equation	Coefficient (R^2)	Half-life ($T_{1/2}$)
Ginseng plant	Fusong	$C_t = 4.9394e^{-0.046t}$	0.9377	13.8
	Huanren	$C_t = 2.7426e^{-0.019t}$	0.8217	28.0
	Baishan	$C_t = 3.9597e^{-0.042t}$	0.9043	13.9
Ginseng soil	Fusong	$C_t = 0.0235e^{-0.019t}$	0.6379	9.8
	Huanren	$C_t = 0.0423e^{-0.028t}$	0.9024	16.0
	Baishan	$C_t = 0.0499e^{-0.014t}$	0.7488	27.4

3.4 Final residues of mandipropamid in ginseng

Based on the design of the field experiment, the final residues of mandipropamid in fresh ginseng, dried ginseng, red ginseng, ginseng plant, and ginseng soil in Fusong

286 County, Jilin Province, Huanren Manchu Autonomous County in Liaoning Province,
287 Baishan City in Jilin Province, Ji'an City in Jilin Province, and Yanbian City in Jilin
288 Province were collected. The results obtained using the above-described methods are
289 shown in Table 4. The residual amounts of mandipropamid were as follows: fresh
290 ginseng, <0.01–0.185; dried ginseng, <0.01–0.265; red ginseng, 0.085–1.544; ginseng
291 plant, 0.075–4.800; and ginseng soil, <0.01–0.014 mg/kg. The final residual amount
292 of mandipropamid in ginseng showed an insignificant decreasing trend with time,
293 which may be affected by the application time, crop varieties, weather, and other
294 factors. The STMR of mandipropamid was < 0.01, 0.019, and 0.295 mg/kg in fresh,
295 dried, and red ginseng, respectively. The HRs of mandipropamid in fresh, dried, and
296 red ginseng were 0.185, 0.265, and 1.544 mg/kg, respectively. At present, China has
297 not established the maximum residue limit (MRL) of mandipropamid in ginseng,
298 partly because of the lack of mandipropamid residue and digestion data available for
299 ginseng. The results of this study provide a basis for the rational use of
300 mandipropamid in ginseng and help establish the MRL of mandipropamid in ginseng.

301 Table 4 Final residues of mandipropamid in ginseng with different application dosages and times

Sample	Does (g a.i./ha)	Times	PHI	Residue data (n = 3) (Mean value, mg/kg)	STMR (mg/kg)	HR (mg/kg)
Fresh ginseng	210.6	1	14	< 0.01(6), 0.034, 0.035, 0.067, 0.185	<0.01	0.185
			21	< 0.01(8), 0.033, 0.033		
			28	< 0.01(7), 0.016, 0.068, 0.069		
Dried ginseng	210.6	1	14	< 0.01, 0.012, 0.012, 0.012, 0.017, 0.019, 0.099, 0.100, 0.222, 0.231	0.019	0.265
			21	0.011, 0.011, 0.015, 0.015, 0.024, 0.024, 0.031, 0.035, 0.157, 0.157		
			28	0.006, 0.007, 0.016, 0.017, 0.018, 0.018, 0.021, 0.021, 0.211, 0.265		
Red ginseng	210.6	1	14	0.085, 0.091, 0.092, 0.100, 0.236, 0.243, 0.255, 0.303, 1.896, 1.901	0.295	1.544
			21	0.164, 0.179, 0.182, 0.289, 0.300, 0.310, 0.380, 0.390, 1.534, 1.544		
			28	0.112, 0.116, 0.133, 0.146, 0.457, 0.462, 0.479, 0.487, 1.432, 1.452		
Ginseng plant	210.6	1	14	0.075, 0.078, 0.223, 0.235, 1.250, 1.255, 1.632, 1.662, 4.643, 4.800	0.729	4.800
			21	0.177, 0.192, 0.500, 0.623, 0.689, 0.694, 0.938, 0.973, 2.005, 2.007		
			28	0.389, 0.466, 0.466, 0.476, 0.578, 0.763, 0.847, 0.872, 1.604, 1.635		
Ginseng soil	210.6	1	14	< 0.01(10)	< 0.01	0.014
			21	<0.01(8), 0.014, 0.014		
			28	<0.01(8), 0.012, 0.012		

302 Note: PHI, Preharvest interval; STMR, supervised trials median residue; HR, highest residue.

3.5 Dietary risk assessment

The chronic dietary exposure risk assessment is based on the Chinese dietary structure, combined with the registered crops and corresponding STMRI values and the daily allowable intake of mandipropamid to calculate the RQ for evaluation. At present, mandipropamid has been registered in China for tomato, pepper, lychee, potato, grape, ginseng, and watermelon. According to the principle of maximum risk, the largest MRL value in the food group was selected for evaluation, and the largest MRL value was selected according to the following priority order: China>CAC>United States>EU>South Korea>Japan>Australia, according to the “National Food Safety Standard Maximum Residue Limits of Pesticides in Food” (National Health Commission of the People's Republic of China, 2019). The ADI of mandipropamid is 0.2 mg/kg bw.

The maximum STMRI value of mandipropamid in fresh, dried, and red ginseng was selected as the reference limit for chronic dietary risk assessment. Ginseng is classified as soy sauce, and the STMRI of mandipropamid in red ginseng is regarded as the residue of soy sauce. The results are shown in Table 5, where the median RQ value is 2.9%, which is far less than 100%, indicating that the chronic dietary risk of mandipropamid use on ginseng is acceptable, and the recommended dose will not cause unacceptable risks to Chinese consumers.

322 Table 5 The Chinese dietary pattern and risk probability of mandipropamid in ginseng

Food classification	Fi (kg)	Residue data (mg/kg)	Sources	NEDI (mg)	ADI (mg)	RQ %
Tubers	0.0495	0.01	China	7.86×10^{-6}		
Dark vegetables	0.0915	3	China	4.36×10^{-3}		
Fruits	0.0457	2	China	1.45×10^{-3}	0.2×63	2.9
Soy sauce	0.009	0.295	STMR	4.21×10^{-5}		
Total	1.0286			5.86×10^{-3}		

323 Note: Fi, Fi, dietary reference intake for a certain kind of food of healthy Chinese people; STMR is the supervised trials median residue of
324 mandipropamid with a pre-harvest interval RQ, chronic dietary exposure risk probability.

4 Conclusion

In this study, an HPLC-MS/MS method was established for the detection of mandipropamid in ginseng, and the degradation dynamics, residue dissipation, and dietary risk assessment of mandipropamid in ginseng were studied under field application. This method has good precision and accuracy. The residues in fresh ginseng, dried ginseng, red ginseng, ginseng plant, and ginseng soil were <0.01–0.185, <0.01–0.265, 0.085–1.544, 0.075–4.800, and <0.01–0.014 mg/kg, respectively. The half-lives of mandipropamid in ginseng plant and soil were 13.8–28.0 and 9.8–27.4 d, respectively. According to the final residual test results, after one application, the chronic dietary risk of mandipropamid in ginseng was an RQ value of 2.9%, which was far less than 100%, indicating that there was no significant potential risk of mandipropamid in ginseng at the recommended dose. This study provides a basis for the rational use of mandipropamid in ginseng, as well as a reference for the establishment of MRL of mandipropamid in ginseng.

Authors Contributions Zhiguang Hou performed interpretation, Writing, Review and Editing.

Xingang Hou performed the data analyses and manuscript preparation and wrote the manuscript.

Liping Wei performed the experiments of field experiments and sample collection section.

Zhanwen Cao performed the experiments of field experiments and sample

346 collection section.

347 Zhou Lu performed the analysis with constructive discussions.

348 Hanju Liu performed the experiments of field experiments and sample collection

349 section.

350 Zhongbin Lu contributed to the conception of the study and provided assistance

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356 **Data availability** All data generated or analyzed during this study are included in this

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358 **Compliance with ethical standards**

359 Ethics approval and consent to participate Not applicable

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361 Competing interests The authors declare that they have no competing interests

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