



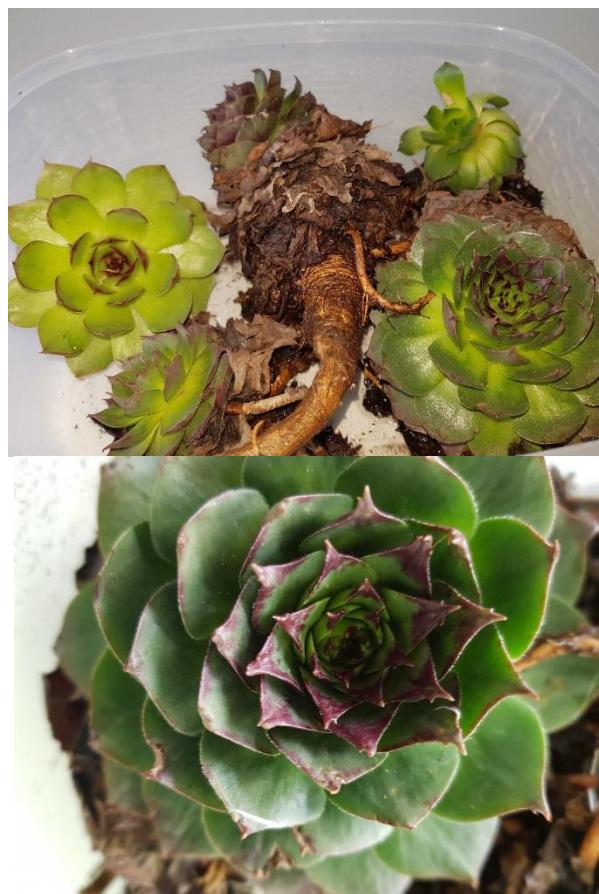
**Figure S1.** *Sempervivum tectorum* L. in the natural habitat (old fortress, Socolari/Ildia - Ciclova Română, Caraș Severin County).



**Figure S2.** *Sempervivum tectorum* L. in the natural habitat.



**Figure S3.** *Sempervivum tectorum* L. (*Sempervivum folium* and *Sempervivum* flower).



**Figure S4.** *Sempervivum tectorum* L. - preparation for processing in the laboratory.



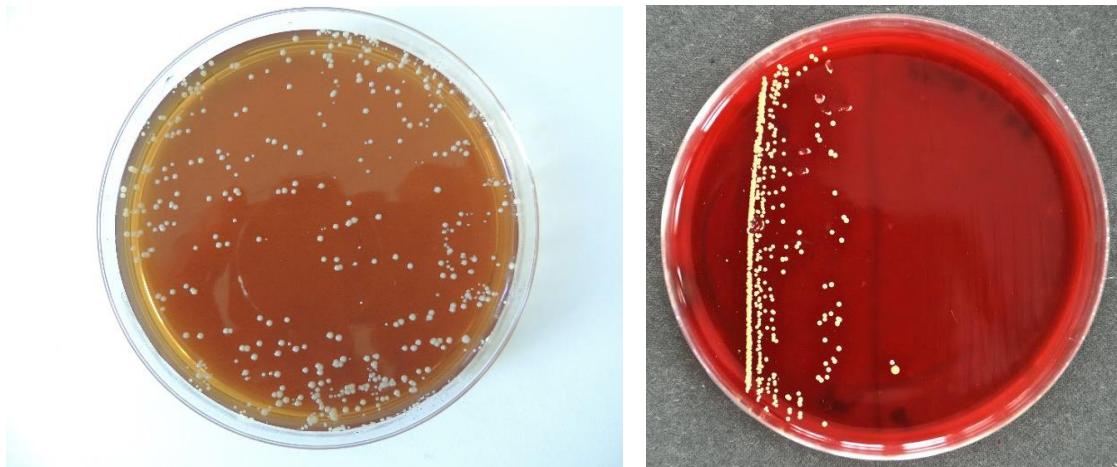
**Figure S5.** *Sempervivum tectorum* L., fresh leaf and lyophilized.



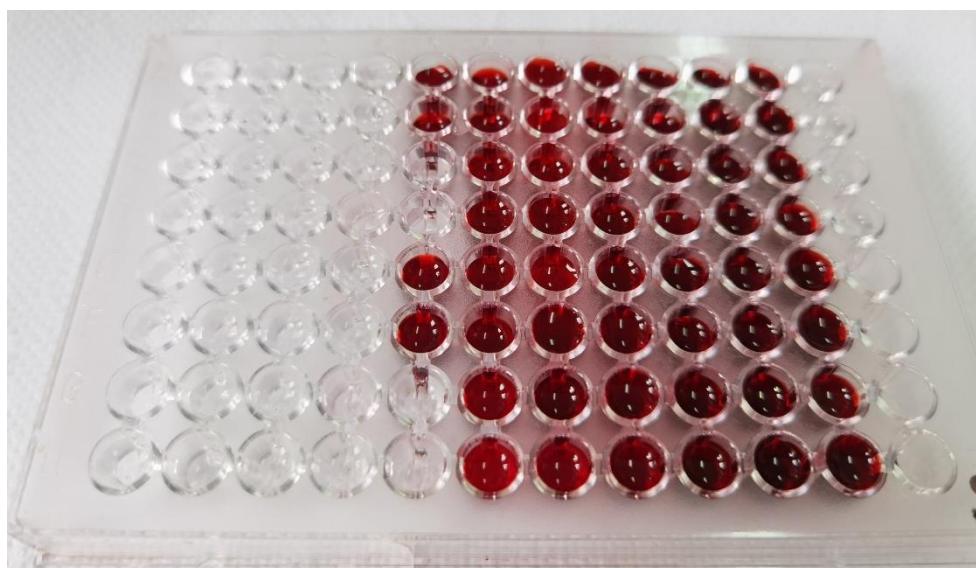
**Figure S6.** Erythematous ceruminous otitis externa in dog.



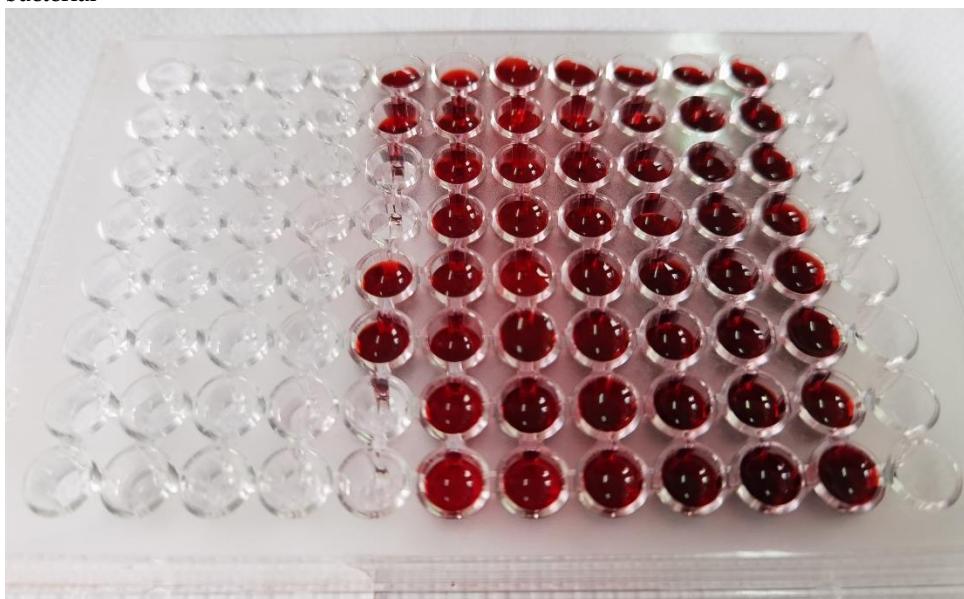
**Figure S7.** *Pseudomonas aeruginosa* (clinical strain).



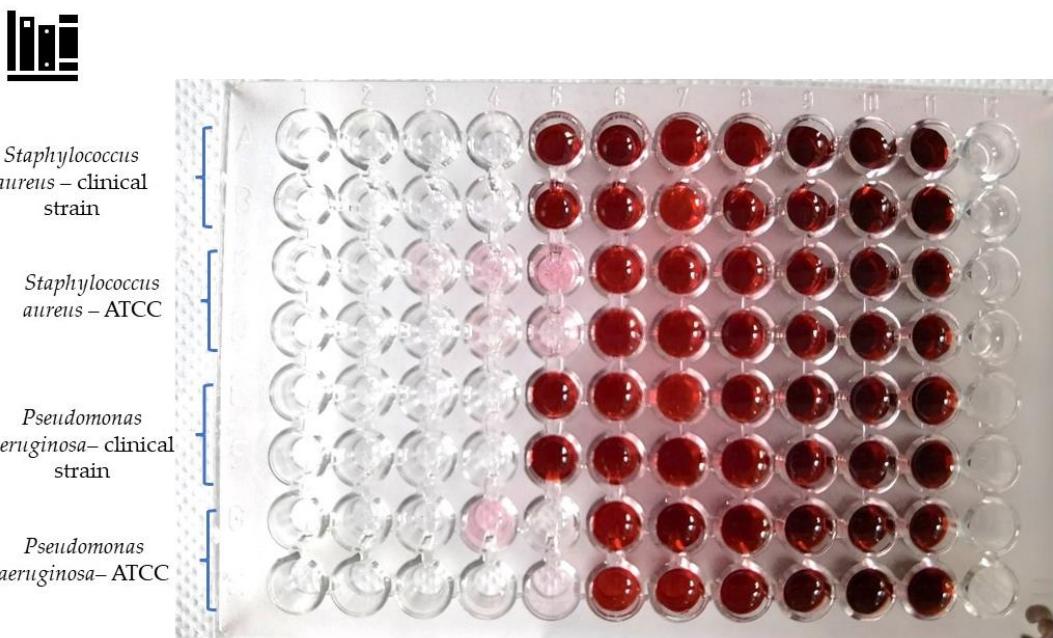
**Figure S8.** *Staphylococcus aureus* (clinical strain).



**Figure S9.** Microtiter plate with varying concentrations of *Sempervivum tectorum L.* extract and tested bacterial strains.



**Figure S10.** Microplate assay for determination of MIC /MBC of *Sempervivum tectorum L.* extract against clinical and standard bacterial strains (*P. aeruginosa* and *S. aureus*).



**Figure S11.** Determination of MIC and MBC , in a 96-well microtiter plate of ethanolic:water extracts of *Sempervivum tectorum L.* against *S. aureus* and *P. aeruginosa* (clinical and standard) strains.

**Table S1.** Total phenol and proanthocyanidin concentration of *S. tectorum L.* extract.

Type of extract	Proanthocyanidin contents (mg PAC/g of extract)	Total phenol contents (mg GA/g of extract)
Ethanolic extract 50% EtOH + 50% H <sub>2</sub> O (leaf)	15.39±0.667 (14.83/15.22/16.13)	126.17±0.334 (125.89/126.08/126.54)

The numbers in parentheses represent the values of the three analyses.

**Table S2.** The median minimal inhibitory (MIC) and minimal bactericidal (MBC) concentrations activity of *S. tectorum L.* extracts (µg/ml).

Bacterial strains	Ethanolic extract 50% EtOH + 50% H <sub>2</sub> O (leaf) - µg/ml			Enrofloxacin (ENR/5µg)	Gentamycin (GM/10µg)
	MIC	MBC	MBC/MIC ratio	MIC	MIC
<i>S. aureus</i> – clinical isolate	23.25	37.23	1.60	0.5	12.5
<i>S. aureus</i> ATCC 25923	20.33	37.29	1.83	0.25	1.125
<i>P. aeruginosa</i> – clinical isolate	24.234	37.30	1.53	0.5	6.25
<i>P. aeruginosa</i> ATCC 27853	20.53	37.02	1.80	0.125	2.5

CLSI, 2020: *P. aeruginosa* breakpoints (µg/ml): enrofloxacin (susceptible - ≤0,5; intermediate – 1-2; resistant ≥4); gentamicin (susceptible - ≤4; intermediate – 4; resistant ≥8); *S. aureus* breakpoints (µg/ml): enrofloxacin (susceptible - ≤0,5; intermediate – 1-2; resistant ≥4); gentamicin (susceptible - ≤4; intermediate – 4; resistant ≥8)

**Table S3.** The minimum inhibitory concentrations of the most effective plant extract against *S. aureus* and *P. aeruginosa* (clinical and standard) strains.

Plant extract	Concentrations of extract (µg/ml)	<i>S. aureus</i> – clinical isolate		<i>S. aureus</i> ATCC 25923		<i>P. aeruginosa</i> – clinical isolate		<i>P. aeruginosa</i> ATCC 27853	
		MIC (µg/ml)	MBC (µg/ml)	MIC (µg/ml)	MBC (µg/ml)	MIC (µg/ml)	MBC (µg/ml)	MIC (µg/ml)	MBC (µg/ml)
	0.5	0	0	0	0	0	0	0	0
	1.0	0	0	0	0	0	0	0	0
	2.0	0	0	1.47± 0.301 (1.15/1.5/1.75)	0	0	0	1.75± 0.25 (1.5/1.75/2)	0
<i>S. tectorum L.</i>	4.0	3.67± 0.381 (3.75/3.25/4)	0	3.53± 0.604 (2.85/3.75/4)	0	3.58± 0.378 (3.15/3.75/3.85)	0	3.66± 0.453 (3.15/3.85/4)	0
<i>L. ethanolic extract</i>	8.0	7.5± 0.661 (6.75/7.75/8)	0	7.11± 0.808 (6.25/7.25/7.8 5)	0	7.7± 0.396 (7.25/7.85/8)	0	7.63± 0.436 (7.15/7.75/8)	0
50% EtOH + 50% H <sub>2</sub> O (leaf)	16.0	14.45± 0.5 (14.75/15.25/15.7 5)	15.93± 0.076 (15.85/15.95/1 6)	15.7± 0.396 (15.25/15.85/1 6)	15.98± 0.028 (16/15.95/16)	15.51± 0.475 (15.05/15.5/16)	15.96± 0.028 (15.95/15.95/1 6)	15.53± 0.604 (14.85/15.75/1 6)	15.75± 0.433 (16/15.25/16)
	32.0	30.05± 1.664 (28.15/30.75/31.2 5)	31.9± 0.132 (31.75/31.95/3 2)	30.83± 1.808 (28.75/31.75/3 2)	31.94± 0.060 (32/31.95/31.8 8)	30.95± 0.818 (30.25/30.75/31.8 5)	31.96± 0.028 (32/31.95/31.9 5)	30.86± 1.125 (29.75/30.85/3 2)	31.58± 0.381 (32/31.25/31.5 
	64.0	63.67± 0.381 (63.25/63.75/64 4)	63.86± 0.132 (63.75/63.85/6 4)	63.36± 0.583 (62.85/63.25/6 4)	63.95± 0.028 (63.85/64/64)	63.43± 0.419 (63.05/63.25/64)	64± 0.00 (64)	63.75± 0.25 (63.5/63.75/64 ))	63.73± 0.421 (64/63.96/63.2 5)

The numbers in parentheses represent the values of the three analyses.

**Table S4.** Phenotypic Antimicrobial Resistance (AMR) profiles of *Staphylococcus aureus* clinical strain.

Antimicrobial	Interpretive Categories and MIC clinical breakpoint ( $\mu\text{g/mL}$ )*			<i>Staphylococcus aureus</i> clinical strain
	Susceptible	Intermediate	Resistant	
Cefoxitin (FOX)	$\leq 4$	-	$\geq 8$	R
Benzylpenicillin (PCG)	$\leq 0.12$	-	$\geq 0.25$	S
Clindamycin (CLI)	$\leq 0.5$	1-2	$\geq 4$	R
Chloramphenicol (CHL)	$\leq 8$	16	$\geq 32$	S
Erythromycin (ERY)	$\leq 0.5$	1-4	$\geq 8$	R
Enrofloxacin (ENR)	$\leq 1$	2	$\geq 4$	S
Fusidic acid (FUS)	$\leq 1$	2	$\geq 4$	S
Gentamicin (GM)	$\leq 4$	8	$\geq 16$	R
Marbofloxacin (MBX)	$\leq 1$	2	$\geq 4$	S
Nitrofurantoin (FT)	$\leq 32$	64	$\geq 128$	S
Oxacillin (OXA)	$\leq 2$	-	$\geq 4$	R
Rifampicin (RIF)	$\leq 1$	2	$\geq 4$	R
Tetracycline (TE)	$\leq 4$	8	$\geq 16$	S
Vancomycin (VAN)	$\leq 2$	4-8	$\geq 16$	R
Trimethoprim/Sulfamethoxazole (SXT)	$\leq 2/38$	-	$\geq 4/76$	R

\*Clinical and Laboratory Standards Institute (CLSI). *Performance Standards for Antimicrobial Susceptibility Testing*. 28th ed. CLSI supplement M100, 2018.

**Table S5.** Phenotypic Antimicrobial Resistance (AMR) profiles of *Pseudomonas aeruginosa* clinical strain.

Antimicrobial	Interpretive Categories and MIC clinical breakpoint ( $\mu\text{g/mL}$ )*			<i>Pseudomonas aeruginosa</i> clinical strain
	Susceptible	Intermediate	Resistant	
Gentamicin (GM)	$\leq 4$	8	$\geq 16$	S
Enrofloxacin (ENR)	$\leq 0.5$	1	$\geq 2$	S
Imipenem (IPM)	$\leq 2$	4	$\geq 8$	R
Meropenem (MEM)	$\leq 2$	4	$\geq 8$	S
Piperacillin-tazobactam (TZP)	$\leq 16/4$	32/4-64/4	$\geq 128/4$	R
Ceftazidime (CAZ)	$\leq 8$	16	$\geq 32$	S
Cefepime (FEP)	$\leq 0.5$	1-2	$\geq 4$	S
Aztreonam (ATM)	$\leq 8$	16	$\geq 32$	S
Azithromycin (AZM)	$\leq 16$	-	$\geq 32$	R
Amikacin (AN)	$\leq 16$	32	$\geq 64$	S
Tobramycin (TM)	$\leq 4$	8	$\geq 16$	R
Polymyxin B (PB)	$\leq 2$	4	$\geq 8$	R
Ciprofloxacin (CIP)	$\leq 0.5$	1	$\geq 2$	S
Doxycycline (DO)	$\leq 4$	8	$\geq 16$	R
Trimethoprim/Sulfamethoxazole (SXT)	$\leq 2/38$	-	$\geq 4/76$	R

\*Clinical and Laboratory Standards Institute (CLSI). *Performance Standards for Antimicrobial Susceptibility Testing*. 28th ed. CLSI supplement M100, 2018.